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NOVEL PROTEINS AND NUCLEIC ACIDS ENCODING SAME

Abstract:

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(71) Applicant *(for all designated States except US)*: CURA-GEN CORPORATION [US/US]; 555 Long Wharf Drive, 11th floor, New Haven, CT 06511 (US).

(72) Inventors; and

(75) Inventors/Applicants *(for US only)*: SMITHSON, Glennda [US/US]; 125 Michael Drive, Guilford, CT 06435 (US). MILLET, Isabelle [FR/US]; 74 Carrington Avenue, Milford, CT 06460 (US). PEYMAN, John, A. [US/US]; 336 West Rock Avenue, New Haven, CT 06515 (US). KEKUDA, Ramesh [IN/US]; 71 Aiken Street, Unit R3, Norwalk, CT 06851 (US). JU, Jingfang [US/US]; 391 Rosebud Lane, Orange, CT 06477 (US). LI, Li [CN/US]; 56 Jerimoth Drive, Branford, CT 06405 (US). GUO, Xiaojia [CN/US]; 713 Robert Frost Drive, Branford, CT 06405 (US). PATTURAJAN, Meera [IN/US]; 45 Harrison Avenue, Apt. 1C, Branford, CT 06405 (US). SPYTEK, Kimberly, A. [US/US]; 28 Court Street #1, New Haven, CT 06511 (US). EDINGER, Shlomit, R. [US/US]; 766 Edgewood Avenue, New Haven, CT 06515 (US). ELLERMAN, Karen [US/US]; 87 Montoya Drive, Branford, CT 06405 (US). MALYANKAR, Uriel, M. [IN/US]; 229 Branford Road, #330, Branford, CT 06405 (US). ORT, Tatiana [—/US]; 85 Viscount Drive, Apartment 61A, Milford, CT 06460 (US). GORMAN, Linda [US/US]; 329 Monticello Drive, Branford, CT 06405 (US). ZERHUSEN, Bryan, D. [US/US]; 337 Monticello Drive, Branford, CT 06405 (US). ANDERSON, David, W. [US/US]; 85 Montoya Drive, Branford, CT 06405 (US). ZHONG, Mei [CA/US]; 45 Harrison Avenue, Apartment 1B, Branford, CT 06405 (US). CATTERTON, Elina [FI/US]; 584, Boston Post Road, Madison, CT 06443 (US). JI, Weizhen [CN/US]; 3 Business Park Drive,

Room 101, Branford, CT 06405 (US). MILLER, Charles, E. [US/US]; 98 Saddle Hill Drive, Guilford, CT 06437 (US). RASTELLI, Luca [IT/US]; 52 Pepperbush Lane, Guilford, CT 06437 (US). STONE, David, J. [US/US]; 223 Whitehorn Drive, Guilford, CN 06437 (US). PENA, Carol, E., A. [US/US]; 604 Orange Street, Number 2, New Haven, CT 06511 (US). SHENOY, Suresh, G. [IN/US]; 15 Millwood Drive, Branford, CT 06405 (US). SHIMKETS, Richard, A. [US/US]; 5 Indian Meadows Drive, Guilford, Connecticut 06437 (US). ROTHEN-BERG, Mark, E. [US/US]; 2 Allen Road, Clinton, CT 06413 (US). LEACH, Martin, D. [GB/US]; 131 Beechwood Drive, Madison, CT 06433 (US). AGEE, Michele, L. [US/US]; 107 Knollwood Road, Wallingford, CT 06492 (US). BERGHS, Constance [NL/US]; 459 Orange Street, New Haven, CT 06511 (US). DIPIPPO, Vincent, A. [US/US]; 156 North Atwater Street, East Haven, CN 06512 (US). EISEN, Andrew, J. [US/US]; 12412 St. James Road, Rockville, MD 20850 (US). GANGOLLI, Esha, A. [IN/US]; 31 Strawberry Hill Road, Madison, CT 06443 (US). RIEGER, Daniel, K. [DE/US]; 10A McKinnel Court, Branford, CT 06405 (US). SPADERNA, Steven, K. [US/US]; 261 Deerfield Drive, Berlin, CT 06037 (US).

(74) Agents: ELRIFI, Ivor, R.; Mintz, Levin, Cohn, Ferris, Glovsky and Popeo, P.C., One Financial Center, Boston, MA 02111 et al. (US).

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NOVEL PROTEINS AND NUCLEIC ACIDS ENCODING SAME

FIELD OF THE INVENTION

The present invention relates to novel polypeptides that are targets of small molecule drugs and that have properties related to stimulation of biochemical or physiological responses in a cell, a tissue, an organ or an organism. More particularly, the novel polypeptides are gene products of novel genes, or are specified biologically active fragments or derivatives thereof. Methods of use encompass diagnostic and prognostic assay procedures as well as methods of treating diverse pathological conditions.

BACKGROUND

Eukaryotic cells are characterized by biochemical and physiological processes which under normal conditions are exquisitely balanced to achieve the preservation and propagation of the cells. When such cells are components of multicellular organisms such

5   as vertebrates, or more particularly organisms such as mammals, the regulation of the biochemical and physiological processes involves intricate signaling pathways. Frequently, such signaling pathways involve extracellular signaling proteins, cellular receptors that bind the signaling proteins and signal transducing components located within the cells.

Signaling proteins may be classified as endocrine effectors, paracrine effectors or

10   autocrine effectors. Endocrine effectors are signaling molecules secreted by a given organ into the circulatory system, which are then transported to a distant target organ or tissue. The target cells include the receptors for the endocrine effector, and when the endocrine effector binds, a signaling cascade is induced. Paracrine effectors involve secreting cells and receptor cells in close proximity to each other, for example two different classes of

15   cells in the same tissue or organ. One class of cells secretes the paracrine effector, which then reaches the second class of cells, for example by diffusion through the extracellular fluid. The second class of cells contains the receptors for the paracrine effector; binding of the effector results in induction of the signaling cascade that elicits the corresponding biochemical or physiological effect. Autocrine effectors are highly analogous to paracrine

20   effectors, except that the same cell type that secretes the autocrine effector also contains the receptor. Thus the autocrine effector binds to receptors on the same cell, or on identical neighboring cells. The binding process then elicits the characteristic biochemical or physiological effect.

Signaling processes may elicit a variety of effects on cells and tissues including by

25   way of nonlimiting example induction of cell or tissue proliferation, suppression of growth or proliferation, induction of differentiation or maturation of a cell or tissue, and suppression of differentiation or maturation of a cell or tissue.

Many pathological conditions involve dysregulation of expression of important effector proteins. In certain classes of pathologies the dysregulation is manifested as

30   diminished or suppressed level of synthesis and secretion of protein effectors. In other classes of pathologies the dysregulation is manifested as increased or up-regulated level of synthesis and secretion of protein effectors. In a clinical setting a subject may be suspected

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of suffering from a condition brought on by altered or mis-regulated levels of a protein effector of interest. Therefore there is a need to assay for the level of the protein effector of interest in a biological sample from such a subject, and to compare the level with that characteristic of a nonpathological condition. There also is a need to provide the protein

5   effector as a product of manufacture. Administration of the effector to a subject in need thereof is useful in treatment of the pathological condition. Accordingly, there is a need for a method of treatment of a pathological condition brought on by a diminished or suppressed levels of the protein effector of interest. In addition, there is a need for a method of treatment of a pathological condition brought on by a increased or up-regulated levels of

10   the protein effector of interest.

Small molecule targets have been implicated in various disease states or pathologies. These targets may be proteins, and particularly enzymatic proteins, which are acted upon by small molecule drugs for the purpose of altering target function and achieving a desired result. Cellular, animal and clinical studies can be performed to

15   elucidate the genetic contribution to the etiology and pathogenesis of conditions in which small molecule targets are implicated in a variety of physiologic, pharmacologic or native states. These studies utilize the core technologies at CuraGen Corporation to look at differential gene expression, protein-protein interactions, large-scale sequencing of expressed genes and the association of genetic variations such as, but not limited to, single

20   nucleotide polymorphisms (SNPs) or splice variants in and between biological samples from experimental and control groups. The goal of such studies is to identify potential avenues for therapeutic intervention in order to prevent, treat the consequences or cure the conditions.

In order to treat diseases, pathologies and other abnormal states or conditions in

25   which a mammalian organism has been diagnosed as being, or as being at risk for becoming, other than in a normal state or condition, it is important to identify new therapeutic agents. Such a procedure includes at least the steps of identifying a target component within an affected tissue or organ, and identifying a candidate therapeutic agent that modulates the functional attributes of the target. The target component may be any

30   biological macromolecule implicated in the disease or pathology. Commonly the target is a polypeptide or protein with specific functional attributes. Other classes of macromolecule may be a nucleic acid, a polysaccharide, a lipid such as a complex lipid or a glycolipid; in addition a target may be a sub-cellular structure or extra-cellular structure that is comprised

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of more than one of these classes of macromolecule. Once such a target has been identified, it may be employed in a screening assay in order to identify favorable candidate therapeutic agents from among a large population of substances or compounds.

In many cases the objective of such screening assays is to identify small molecule
5   candidates; this is commonly approached by the use of combinatorial methodologies to develop the population of substances to be tested. The implementation of high throughput screening methodologies is advantageous when working with large, combinatorial libraries of compounds.

SUMMARY OF THE INVENTION

10   The invention includes nucleic acid sequences and the novel polypeptides they encode. The novel nucleic acids and polypeptides are referred to herein as NOVX, or NOV1, NOV2, NOV3, *etc.*, nucleic acids and polypeptides. These nucleic acids and polypeptides, as well as derivatives, homologs, analogs and fragments thereof, will hereinafter be collectively designated as "NOVX" nucleic acid, which represents the
15   nucleotide sequence selected from the group consisting of SEQ ID NO: 2n-1, wherein n is an integer between 1 and 124, or polypeptide sequences, which represents the group consisting of SEQ ID NO: 2n, wherein n is an integer between 1 and 124.

In one aspect, the invention provides an isolated polypeptide comprising a mature form of a NOVX amino acid. One example is a variant of a mature form of a NOVX
20   amino acid sequence, wherein any amino acid in the mature form is changed to a different amino acid, provided that no more than 15% of the amino acid residues in the sequence of the mature form are so changed. The amino acid can be, for example, a NOVX amino acid sequence or a variant of a NOVX amino acid sequence, wherein any amino acid specified in the chosen sequence is changed to a different amino acid, provided that no more than
25   15% of the amino acid residues in the sequence are so changed. The invention also includes fragments of any of these. In another aspect, the invention also includes an isolated nucleic acid that encodes a NOVX polypeptide, or a fragment, homolog, analog or derivative thereof.

Also included in the invention is a NOVX polypeptide that is a naturally occurring
30   allelic variant of a NOVX sequence. In one embodiment, the allelic variant includes an amino acid sequence that is the translation of a nucleic acid sequence differing by a single nucleotide from a NOVX nucleic acid sequence. In another embodiment, the NOVX

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polypeptide is a variant polypeptide described therein, wherein any amino acid specified in the chosen sequence is changed to provide a conservative substitution. In one embodiment, the invention discloses a method for determining the presence or amount of the NOVX polypeptide in a sample. The method involves the steps of: providing a sample;

5   introducing the sample to an antibody that binds immunospecifically to the polypeptide; and determining the presence or amount of antibody bound to the NOVX polypeptide, thereby determining the presence or amount of the NOVX polypeptide in the sample. In another embodiment, the invention provides a method for determining the presence of or predisposition to a disease associated with altered levels of a NOVX polypeptide in a

10   mammalian subject. This method involves the steps of: measuring the level of expression of the polypeptide in a sample from the first mammalian subject; and comparing the amount of the polypeptide in the sample of the first step to the amount of the polypeptide present in a control sample from a second mammalian subject known not to have, or not to be predisposed to, the disease, wherein an alteration in the expression level of the

15   polypeptide in the first subject as compared to the control sample indicates the presence of or predisposition to the disease.

       In a further embodiment, the invention includes a method of identifying an agent that binds to a NOVX polypeptide. This method involves the steps of: introducing the polypeptide to the agent; and determining whether the agent binds to the polypeptide. In

20   various embodiments, the agent is a cellular receptor or a downstream effector.

       In another aspect, the invention provides a method for identifying a potential therapeutic agent for use in treatment of a pathology, wherein the pathology is related to aberrant expression or aberrant physiological interactions of a NOVX polypeptide. The method involves the steps of: providing a cell expressing the NOVX polypeptide and

25   having a property or function ascribable to the polypeptide; contacting the cell with a composition comprising a candidate substance; and determining whether the substance alters the property or function ascribable to the polypeptide; whereby, if an alteration observed in the presence of the substance is not observed when the cell is contacted with a composition devoid of the substance, the substance is identified as a potential therapeutic

30   agent. In another aspect, the invention describes a method for screening for a modulator of activity or of latency or predisposition to a pathology associated with the NOVX polypeptide. This method involves the following steps: administering a test compound to a test animal at increased risk for a pathology associated with the NOVX polypeptide,

wherein the test animal recombinantly expresses the NOVX polypeptide. This method involves the steps of measuring the activity of the NOVX polypeptide in the test animal after administering the compound of step; and comparing the activity of the protein in the test animal with the activity of the NOVX polypeptide in a control animal not administered

5   the polypeptide, wherein a change in the activity of the NOVX polypeptide in the test animal relative to the control animal indicates the test compound is a modulator of latency of, or predisposition to, a pathology associated with the NOVX polypeptide. In one embodiment, the test animal is a recombinant test animal that expresses a test protein transgene or expresses the transgene under the control of a promoter at an increased level

10   relative to a wild-type test animal, and wherein the promoter is not the native gene promoter of the transgene. In another aspect, the invention includes a method for modulating the activity of the NOVX polypeptide, the method comprising introducing a cell sample expressing the NOVX polypeptide with a compound that binds to the polypeptide in an amount sufficient to modulate the activity of the polypeptide.

15   The invention also includes an isolated nucleic acid that encodes a NOVX polypeptide, or a fragment, homolog, analog or derivative thereof. In a preferred embodiment, the nucleic acid molecule comprises the nucleotide sequence of a naturally occurring allelic nucleic acid variant. In another embodiment, the nucleic acid encodes a variant polypeptide, wherein the variant polypeptide has the polypeptide sequence of a

20   naturally occurring polypeptide variant. In another embodiment, the nucleic acid molecule differs by a single nucleotide from a NOVX nucleic acid sequence. In one embodiment, the NOVX nucleic acid molecule hybridizes under stringent conditions to the nucleotide sequence selected from the group consisting of SEQ ID NO: 2n-1, wherein n is an integer between 1 and 124, or a complement of the nucleotide sequence. In another aspect, the

25   invention provides a vector or a cell expressing a NOVX nucleotide sequence.

In one embodiment, the invention discloses a method for modulating the activity of a NOVX polypeptide. The method includes the steps of: introducing a cell sample expressing the NOVX polypeptide with a compound that binds to the polypeptide in an amount sufficient to modulate the activity of the polypeptide. In another embodiment, the

30   invention includes an isolated NOVX nucleic acid molecule comprising a nucleic acid sequence encoding a polypeptide comprising a NOVX amino acid sequence or a variant of a mature form of the NOVX amino acid sequence, wherein any amino acid in the mature form of the chosen sequence is changed to a different amino acid, provided that no more

than 15% of the amino acid residues in the sequence of the mature form are so changed. In another embodiment, the invention includes an amino acid sequence that is a variant of the NOVX amino acid sequence, in which any amino acid specified in the chosen sequence is changed to a different amino acid, provided that no more than 15% of the amino acid

5   residues in the sequence are so changed.

In one embodiment, the invention discloses a NOVX nucleic acid fragment encoding at least a portion of a NOVX polypeptide or any variant of the polypeptide, wherein any amino acid of the chosen sequence is changed to a different amino acid, provided that no more than 10% of the amino acid residues in the sequence are so changed.

10   In another embodiment, the invention includes the complement of any of the NOVX nucleic acid molecules or a naturally occurring allelic nucleic acid variant. In another embodiment, the invention discloses a NOVX nucleic acid molecule that encodes a variant polypeptide, wherein the variant polypeptide has the polypeptide sequence of a naturally occurring polypeptide variant. In another embodiment, the invention discloses a NOVX

15   nucleic acid, wherein the nucleic acid molecule differs by a single nucleotide from a NOVX nucleic acid sequence.

In another aspect, the invention includes a NOVX nucleic acid, wherein one or more nucleotides in the NOVX nucleotide sequence is changed to a different nucleotide provided that no more than 15% of the nucleotides are so changed. In one embodiment, the

20   invention discloses a nucleic acid fragment of the NOVX nucleotide sequence and a nucleic acid fragment wherein one or more nucleotides in the NOVX nucleotide sequence is changed from that selected from the group consisting of the chosen sequence to a different nucleotide provided that no more than 15% of the nucleotides are so changed. In another embodiment, the invention includes a nucleic acid molecule wherein the nucleic

25   acid molecule hybridizes under stringent conditions to a NOVX nucleotide sequence or a complement of the NOVX nucleotide sequence. In one embodiment, the invention includes a nucleic acid molecule, wherein the sequence is changed such that no more than 15% of the nucleotides in the coding sequence differ from the NOVX nucleotide sequence or a fragment thereof.

30   In a further aspect, the invention includes a method for determining the presence or amount of the NOVX nucleic acid in a sample. The method involves the steps of: providing the sample; introducing the sample to a probe that binds to the nucleic acid molecule; and determining the presence or amount of the probe bound to the NOVX

nucleic acid molecule, thereby determining the presence or amount of the NOVX nucleic acid molecule in the sample. In one embodiment, the presence or amount of the nucleic acid molecule is used as a marker for cell or tissue type.

5　　　　　In another aspect, the invention discloses a method for determining the presence of or predisposition to a disease associated with altered levels of the NOVX nucleic acid molecule of in a first mammalian subject. The method involves the steps of: measuring the amount of NOVX nucleic acid in a sample from the first mammalian subject; and comparing the amount of the nucleic acid in the sample of step (a) to the amount of NOVX nucleic acid present in a control sample from a second mammalian subject known not to

10　　have or not be predisposed to, the disease; wherein an alteration in the level of the nucleic acid in the first subject as compared to the control sample indicates the presence of or predisposition to the disease.

　　　　　Unless otherwise defined, all technical and scientific terms used herein have the same meaning as commonly understood by one of ordinary skill in the art to which this

15　　invention belongs. Although methods and materials similar or equivalent to those described herein can be used in the practice or testing of the present invention, suitable methods and materials are described below. All publications, patent applications, patents, and other references mentioned herein are incorporated by reference in their entirety. In the case of conflict, the present specification, including definitions, will control. In

20　　addition, the materials, methods, and examples are illustrative only and not intended to be limiting.

　　　　　Other features and advantages of the invention will be apparent from the following detailed description and claims.

DETAILED DESCRIPTION OF THE INVENTION

25　　　　　The present invention provides novel nucleotides and polypeptides encoded thereby. Included in the invention are the novel nucleic acid sequences, their encoded polypeptides, antibodies, and other related compounds. The sequences are collectively referred to herein as "NOVX nucleic acids" or "NOVX polynucleotides" and the corresponding encoded polypeptides are referred to as "NOVX polypeptides" or "NOVX

30　　proteins." Unless indicated otherwise, "NOVX" is meant to refer to any of the novel sequences disclosed herein. Table A provides a summary of the NOVX nucleic acids and their encoded polypeptides.

TABLE A. Sequences and Corresponding SEQ ID Numbers

NOVX Assignment	Internal Identification	SEQ ID NO (nucleic acid)	SEQ ID NO (amino acid)	Homology
1a	CG106764-01	1	2	Citron Kinase
1b	268667493	3	4	RHO/RAC-Interacting Citron Kinase
1c	268667539	5	6	RHO/RAC-Interacting Citron Kinase
1d	268667543	7	8	RHO/RAC-Interacting Citron Kinase
1e	268667555	9	10	RHO/RAC-Interacting Citron Kinase
1f	268667574	11	12	RHO/RAC-Interacting Citron Kinase
1g	CG106764-02	13	14	RHO/RAC-Interacting Citron Kinase
2a	CG117662-01	15	16	Renal Renin Precursor
2b	CG117662-02	17	18	Renal Renin Precursor
3a	CG118051-01	19	20	Aldehyde Dehydrogenase
3b	CG118051-02	21	22	Aldehyde Dehydrogenase
3c	CG118051-03	23	24	Aldehyde Dehydrogenase
4a	CG120277-01	25	26	Aldehyde Dehydrogenase-3
4b	CG120277-02	27	28	Aldehyde Dehydrogenase-3
5a	CG140468-01	29	30	Serine/Threonine-Protein Kinase PAK 1
5b	CG140468-02	31	32	Serine/Threonine-Protein Kinase PAK 1
6a	CG142182-01	33	34	Ubiquitin Carboxyl-terminal Hydrolase 15
7a	CG142564-01	35	36	Carnitine O-Palmitoyltransferase I
8a	CG142797-01	37	38	Cathepsin L
9a	CG143216-01	39	40	Laminin Gamma 3 Chain Precursor
10a	CG143787-01	41	42	Disintegrin Protease
10b	278889162	43	44	Disintegrin Protease
10c	278689868	45	46	Disintegrin Protease
11a	CG144112-01	47	48	NEUROPSIN PRECURSOR like homo sapiens
11b	CG144112-04	49	50	Neuropsin Precursor
11c	255501898	51	52	Neuropsin Precursor
11d	255612524	53	54	Neuropsin Precursor
11e	255612566	55	56	Neuropsin Precursor
11f	306434072	57	58	Neuropsin Precursor
11g	CG144112-02	59	60	Neuropsin Precursor
11h	CG144112-03	61	62	Neuropsin Precursor
12a	CG144497-01	63	64	Adenylosuccinate Synthetase Muscle Isozyme
13a	CG144686-01	65	66	Mast Cell Carboxypeptidase A Precursor
13b	278690008	67	68	Mast Cell Carboxypeptidase A Precursor
13c	278690035	69	70	Mast Cell Carboxypeptidase A Precursor
13d	CG144686-02	71	72	Mast Cell Carboxypeptidase A Precursor
14a	CG144906-01	73	74	Testisin Precursor
14b	CG144906-02	75	76	Testisin Precursor
15a	CG144997-01	77	78	RNase H I
15b	278693648	79	80	RNase H I
15c	278480974	81	82	RNase H I
15d	278498047	83	84	RNase H I

15e	CG144997-02	85	86	RNase H1
16a	CG145494-01	87	88	PRESTIN
17a	CG145722-01	89	90	WEE1
18a	CG145754-01	91	92	Kallikrein 7 Precursor
18b	CG145754-03	93	94	Kallikrein 7 Precursor
18c	CG145754-02	95	96	Kallikrein 7 Precursor
18d	252718128	97	98	Kallikrein
18e	252718152	99	100	Kallikrein
18f	247856668	101	102	Kallikrein 7 Precursor
18g	247856705	103	104	Kallikrein 7 Precursor
19a	CG146279-01	105	106	Novel Potassium Channel Subfamily K Member 10 (TREK-2)
20a	CG146374-01	107	108	Glycogen Branching Enzyme
21a	CG146403-01	109	110	Diacylglycerol Acyltransferase 2
22a	CG146513-01	111	112	Diacylglycerol Acyltransferase 2
23a	CG146522-01	113	114	Diacylglycerol Acyltransferase 2
24a	CG146531-01	115	116	Diacylglycerol Acyltransferase 2
25a	CG147274-01	117	118	Protease
26a	CG147351-01	119	120	Testis-Development Related NYD-SP27
27a	CG147419-01	121	122	Glutamine:Fructose-6-Phosphate Amidotransferase 1 Muscle Isoform
28a	CG148102-01	123	124	Carnitine O-Palmitoyltransferase
28b	CG148102-02	125	126	Carnitine O-Palmitoyltransferase
29a	CG148431-01	127	128	Class II Aminotransferase
29b	CG148431-02	129	130	Class II Aminotransferase
30a	CG148888-01	131	132	GALNAC 4-Sulfotransferase
31a	CG149008-01	133	134	Sodium/Hydrogen Exchanger
32a	CG149350-01	135	136	Vacuolar ATP Synthase Subunit F
32b	CG149350-02	137	138	Vacuolar ATP Synthase Subunit F
33a	CG149463-01	139	140	Serine/Threonine-Protein Kinase SGK
34a	CG149536-01	141	142	Protein-Tyrosine Phosphatase, Non-Receptor Type 2
35a	CG149964-01	143	144	Brain Mitochondrial Carrier Protein-1
35b	309326356	145	146	Brain Mitochondrial Carrier Protein-1
35c	309326444	147	148	Brain Mitochondrial Carrier Protein-1
35d	309326473	149	150	Brain Mitochondrial Carrier Protein-1
35e	CG149964-02	151	152	Brain Mitochondrial Carrier Protein-1
36a	CG150306-01	153	154	Dual Specificity Protein Phosphatase 5
37a	CG150510-01	155	156	Human Alpha-2,3-Sialyltransferase
38a	CG150704-01	157	158	Testis ecto-ADP-Ribosyltransferase Precursor
39a	CG150799-01	159	160	MASS1
39b	CG150799-02	161	162	MASS1
39c	CG150799-03	163	164	MASS1
39d	CG150799-01	165	166	MASS1
40a	CG151014-01	167	168	Metabotropic Glutamate Receptor 3
40b	CG151014-02	169	170	Metabotropic Glutamate Receptor 3
40c	CG151014-03	171	172	Metabotropic Glutamate Receptor 3
41a	CG151297-01	173	174	Calmodulin-Dependent Phosphodiesterase
41b	CG151297-02	175	176	Calmodulin-Dependent Phosphodiesterase

42a	CG151822-01	177	178	Prenylcysteine Carboxyl Methyltransferase
42b	CG151822-02	179	180	Prenylcysteine Carboxyl Methyltransferase
43a	CG152256-01	181	182	Phosphatidylserine Synthase
44a	CG171804-01	183	184	N-Acetylgalactosaminide Alpha 2, 6-Sialyltransferase
45a	CG171841-01	185	186	Iron-Containing Alcohol Dehydrogenase
46a	CG173017-01	187	188	Retinoic Acid Receptor RXR-Beta
47a	CG173347-01	189	190	Serum Paraoxonase/Arylesterase 3
48a	CG56234-01	191	192	Phosphoenolpyruvate Carboxykinase 2 (PCK2)
48b	CG56234-02	193	194	Phosphoenolpyruvate Carboxykinase 2 (PCK2)
49a	CG56836-01	195	196	Cathepsin B
49b	CG56836-02	197	198	Cathepsin B
49c	CG56836-03	199	200	Cathepsin B
49d	CG56836-04	201	202	Cathepsin B
49e	247856403	203	204	Cathepsin B
49f	247856434	205	206	Cathepsin B
49g	247856497	207	208	Cathepsin B
49h	247856493	209	210	Cathepsin B
49i	247856574	211	212	Cathepsin B
49j	247856545	213	214	Cathepsin B
49k	275480714	215	216	Cathepsin B
50a	CG57284-01	217	218	RAS-Related Protein RAB-5C
50b	CG57284-03	219	220	RAS-Related Protein RAB-5C
50c	CG57284-02	221	222	RAS-Related Protein RAB-5C
51a	CG57308-01	223	224	Sulfonylurea Receptor 1
51b	CG57308-02	225	226	Sulfonylurea Receptor 1
52a	CG93659-01	227	228	Mitogen-Activated Protein Kinase Kinase Kinase 8
52b	CG93659-03	229	230	Mitogen-Activated Protein Kinase Kinase Kinase 8
52c	CG93659-02	231	232	Mitogen-Activated Protein Kinase Kinase Kinase 8
53a	CG94521-01	233	234	Cytoplasmic Glycerol-3-Phosphate Dehydrogenase [NAD+]
53b	CG94521-03	235	236	Cytoplasmic Glycerol-3-Phosphate Dehydrogenase [NAD+]
53c	CG94521-02	237	238	Cytoplasmic Glycerol-3-Phosphate Dehydrogenase [NAD+]
54a	CG96613-01	239	240	Pyruvate Dehydrogenase Kinase (PDK1)
54b	CG96613-03	241	242	Pyruvate Dehydrogenase Kinase (PDK1)
54c	CG96613-02	243	244	Pyruvate Dehydrogenase Kinase (PDK1)
55a	CG96736-01	245	246	Neutral Amino Acid Transporter B
55b	CG96736-02	247	248	Neutral Amino Acid Transporter B

Table A indicates the homology of NOVX polypeptides to known protein families.

5   Thus, the nucleic acids and polypeptides, antibodies and related compounds according to

the invention corresponding to a NOVX as identified in column 1 of Table A will be useful in therapeutic and diagnostic applications implicated in, for example, pathologies and disorders associated with the known protein families identified in column 5 of Table A.

Pathologies, diseases, disorders and condition and the like that are associated with

5   NOVX sequences include, but are not limited to: *e.g.*, cardiomyopathy, atherosclerosis, hypertension, congenital heart defects, aortic stenosis, atrial septal defect (ASD), atrioventricular (A-V) canal defect, ductus arteriosus, pulmonary stenosis, subaortic stenosis, ventricular septal defect (VSD), valve diseases, tuberous sclerosis, scleroderma, obesity, metabolic disturbances associated with obesity, transplantation,

10   adrenoleukodystrophy, congenital adrenal hyperplasia, prostate cancer, diabetes, metabolic disorders, neoplasm; adenocarcinoma, lymphoma, uterus cancer, fertility, hemophilia, hypercoagulation, idiopathic thrombocytopenic purpura, immunodeficiencies, graft versus host disease, AIDS, bronchial asthma, Crohn's disease; multiple sclerosis, treatment of Albright Hereditary Ostoeodystrophy, infectious disease, anorexia, cancer-associated

15   cachexia, cancer, neurodegenerative disorders, Alzheimer's Disease, Parkinson's Disorder, immune disorders, hematopoietic disorders, and the various dyslipidemias, the metabolic syndrome X and wasting disorders associated with chronic diseases and various cancers, as well as conditions such as transplantation and fertility.

NOVX nucleic acids and their encoded polypeptides are useful in a variety of

20   applications and contexts. The various NOVX nucleic acids and polypeptides according to the invention are useful as novel members of the protein families according to the presence of domains and sequence relatedness to previously described proteins. Additionally, NOVX nucleic acids and polypeptides can also be used to identify proteins that are members of the family to which the NOVX polypeptides belong.

25   Consistent with other known members of the family of proteins, identified in column 5 of Table A, the NOVX polypeptides of the present invention show homology to, and contain domains that are characteristic of, other members of such protein families. Details of the sequence relatedness and domain analysis for each NOVX are presented in Example A.

30   The NOVX nucleic acids and polypeptides can also be used to screen for molecules, which inhibit or enhance NOVX activity or function. Specifically, the nucleic acids and polypeptides according to the invention may be used as targets for the identification of

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small molecules that modulate or inhibit diseases associated with the protein families listed in Table A.

The NOVX nucleic acids and polypeptides are also useful for detecting specific cell types. Details of the expression analysis for each NOVX are presented in Example C. Accordingly, the NOVX nucleic acids, polypeptides, antibodies and related compounds according to the invention will have diagnostic and therapeutic applications in the detection of a variety of diseases with differential expression in normal vs. diseased tissues, *e.g.* detection of a variety of cancers.

Additional utilities for NOVX nucleic acids and polypeptides according to the invention are disclosed herein.

NOVX clones

NOVX nucleic acids and their encoded polypeptides are useful in a variety of applications and contexts. The various NOVX nucleic acids and polypeptides according to the invention are useful as novel members of the protein families according to the presence of domains and sequence relatedness to previously described proteins. Additionally, NOVX nucleic acids and polypeptides can also be used to identify proteins that are members of the family to which the NOVX polypeptides belong.

The NOVX genes and their corresponding encoded proteins are useful for preventing, treating or ameliorating medical conditions, *e.g.*, by protein or gene therapy. Pathological conditions can be diagnosed by determining the amount of the new protein in a sample or by determining the presence of mutations in the new genes. Specific uses are described for each of the NOVX genes, based on the tissues in which they are most highly expressed. Uses include developing products for the diagnosis or treatment of a variety of diseases and disorders.

The NOVX nucleic acids and proteins of the invention are useful in potential diagnostic and therapeutic applications and as a research tool. These include serving as a specific or selective nucleic acid or protein diagnostic and/or prognostic marker, wherein the presence or amount of the nucleic acid or the protein are to be assessed, as well as potential therapeutic applications such as the following: (i) a protein therapeutic, (ii) a small molecule drug target, (iii) an antibody target (therapeutic, diagnostic, drug targeting/cytotoxic antibody), (iv) a nucleic acid useful in gene therapy (gene delivery/gene ablation), and (v) a composition promoting tissue regeneration *in vitro* and *in vivo* (vi) a biological defense weapon.

In one specific embodiment, the invention includes an isolated polypeptide comprising an amino acid sequence selected from the group consisting of: (a) a mature form of the amino acid sequence selected from the group consisting of SEQ ID NO: 2n, wherein n is an integer between 1 and 124; (b) a variant of a mature form of the amino acid

5   sequence selected from the group consisting of SEQ ID NO: 2n, wherein n is an integer between 1 and 124, wherein any amino acid in the mature form is changed to a different amino acid, provided that no more than 15% of the amino acid residues in the sequence of the mature form are so changed; (c) an amino acid sequence selected from the group consisting of SEQ ID NO: 2n, wherein n is an integer between 1 and 124; (d) a variant of

10   the amino acid sequence selected from the group consisting of SEQ ID NO:2n, wherein n is an integer between 1 and 124 wherein any amino acid specified in the chosen sequence is changed to a different amino acid, provided that no more than 15% of the amino acid residues in the sequence are so changed; and (e) a fragment of any of (a) through (d).

In another specific embodiment, the invention includes an isolated nucleic acid

15   molecule comprising a nucleic acid sequence encoding a polypeptide comprising an amino acid sequence selected from the group consisting of: (a) a mature form of the amino acid sequence given SEQ ID NO: 2n, wherein n is an integer between 1 and 124; (b) a variant of a mature form of the amino acid sequence selected from the group consisting of SEQ ID NO: 2n, wherein n is an integer between 1 and 124 wherein any amino acid in the mature

20   form of the chosen sequence is changed to a different amino acid, provided that no more than 15% of the amino acid residues in the sequence of the mature form are so changed; (c) the amino acid sequence selected from the group consisting of SEQ ID NO: 2n, wherein n is an integer between 1 and 124; (d) a variant of the amino acid sequence selected from the group consisting of SEQ ID NO: 2n, wherein n is an integer between 1 and 124, in which

25   any amino acid specified in the chosen sequence is changed to a different amino acid, provided that no more than 15% of the amino acid residues in the sequence are so changed; (e) a nucleic acid fragment encoding at least a portion of a polypeptide comprising the amino acid sequence selected from the group consisting of SEQ ID NO: 2n, wherein n is an integer between 1 and 124 or any variant of said polypeptide wherein any amino acid of the

30   chosen sequence is changed to a different amino acid, provided that no more than 10% of the amino acid residues in the sequence are so changed; and (f) the complement of any of said nucleic acid molecules.

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In yet another specific embodiment, the invention includes an isolated nucleic acid molecule, wherein said nucleic acid molecule comprises a nucleotide sequence selected from the group consisting of: (a) the nucleotide sequence selected from the group consisting of SEQ ID NO: 2n-1, wherein n is an integer between 1 and 124; (b) a

5   nucleotide sequence wherein one or more nucleotides in the nucleotide sequence selected from the group consisting of SEQ ID NO: 2n-1, wherein n is an integer between 1 and 124 is changed from that selected from the group consisting of the chosen sequence to a different nucleotide provided that no more than 15% of the nucleotides are so changed; (c) a nucleic acid fragment of the sequence selected from the group consisting of SEQ ID

10   NO: 2n-1, wherein n is an integer between 1 and 124; and (d) a nucleic acid fragment wherein one or more nucleotides in the nucleotide sequence selected from the group consisting of SEQ ID NO: 2n-1, wherein n is an integer between 1 and 124 is changed from that selected from the group consisting of the chosen sequence to a different nucleotide provided that no more than 15% of the nucleotides are so changed.

15   **NOVX Nucleic Acids and Polypeptides**

One aspect of the invention pertains to isolated nucleic acid molecules that encode NOVX polypeptides or biologically active portions thereof. Also included in the invention are nucleic acid fragments sufficient for use as hybridization probes to identify NOVX-encoding nucleic acids (*e.g.*, NOVX mRNAs) and fragments for use as PCR

20   primers for the amplification and/or mutation of NOVX nucleic acid molecules. As used herein, the term "nucleic acid molecule" is intended to include DNA molecules (*e.g.*, cDNA or genomic DNA), RNA molecules (*e.g.*, mRNA), analogs of the DNA or RNA generated using nucleotide analogs, and derivatives, fragments and homologs thereof. The nucleic acid molecule may be single-stranded or double-stranded, but preferably is

25   comprised double-stranded DNA.

A NOVX nucleic acid can encode a mature NOVX polypeptide. As used herein, a "mature" form of a polypeptide or protein disclosed in the present invention is the product of a naturally occurring polypeptide or precursor form or proprotein. The naturally occurring polypeptide, precursor or proprotein includes, by way of nonlimiting example,

30   the full-length gene product encoded by the corresponding gene. Alternatively, it may be defined as the polypeptide, precursor or proprotein encoded by an ORF described herein. The product "mature" form arises, by way of nonlimiting example, as a result of one or more naturally occurring processing steps that may take place within the cell (*e.g.*, host

cell) in which the gene product arises. Examples of such processing steps leading to a "mature" form of a polypeptide or protein include the cleavage of the N-terminal methionine residue encoded by the initiation codon of an ORF, or the proteolytic cleavage of a signal peptide or leader sequence. Thus a mature form arising from a precursor

5   polypeptide or protein that has residues 1 to N, where residue 1 is the N-terminal methionine, would have residues 2 through N remaining after removal of the N-terminal methionine. Alternatively, a mature form arising from a precursor polypeptide or protein having residues 1 to N, in which an N-terminal signal sequence from residue 1 to residue M is cleaved, would have the residues from residue M+1 to residue N remaining. Further as

10 used herein, a "mature" form of a polypeptide or protein may arise from a step of post-translational modification other than a proteolytic cleavage event. Such additional processes include, by way of non-limiting example, glycosylation, myristylation or phosphorylation. In general, a mature polypeptide or protein may result from the operation of only one of these processes, or a combination of any of them.

15   The term "probe", as utilized herein, refers to nucleic acid sequences of variable length, preferably between at least about 10 nucleotides (nt), about 100 nt, or as many as approximately, *e.g.*, 6,000 nt, depending upon the specific use. Probes are used in the detection of identical, similar, or complementary nucleic acid sequences. Longer length probes are generally obtained from a natural or recombinant source, are highly specific, and

20 much slower to hybridize than shorter-length oligomer probes. Probes may be single-stranded or double-stranded and designed to have specificity in PCR, membrane-based hybridization technologies, or ELISA-like technologies.

The term "isolated" nucleic acid molecule, as used herein, is a nucleic acid that is separated from other nucleic acid molecules which are present in the natural source of the

25 nucleic acid. Preferably, an "isolated" nucleic acid is free of sequences which naturally flank the nucleic acid (*i.e.*, sequences located at the 5'- and 3'-termini of the nucleic acid) in the genomic DNA of the organism from which the nucleic acid is derived. For example, in various embodiments, the isolated NOVX nucleic acid molecules can contain less than about 5 kb, 4 kb, 3 kb, 2 kb, 1 kb, 0.5 kb or 0.1 kb of nucleotide sequences which naturally

30 flank the nucleic acid molecule in genomic DNA of the cell/tissue from which the nucleic acid is derived (*e.g.*, brain, heart, liver, spleen, *etc.*). Moreover, an "isolated" nucleic acid molecule, such as a cDNA molecule, can be substantially free of other cellular material, or culture medium, or of chemical precursors or other chemicals.

A nucleic acid molecule of the invention, *e.g.*, a nucleic acid molecule having the nucleotide sequence of SEQ ID NO:$2n$-1, wherein n is an integer between 1 and 124, or a complement of this nucleotide sequence, can be isolated using standard molecular biology techniques and the sequence information provided herein. Using all or a portion of the

5   nucleic acid sequence of SEQ ID NO:$2n$-1, wherein n is an integer between 1 and 124, as a hybridization probe, NOVX molecules can be isolated using standard hybridization and cloning techniques (*e.g.*, as described in Sambrook, *et al.*, (eds.), MOLECULAR CLONING: A LABORATORY MANUAL 2nd Ed., Cold Spring Harbor Laboratory Press, Cold Spring Harbor, NY, 1989; and Ausubel, *et al.*, (eds.), CURRENT PROTOCOLS IN MOLECULAR BIOLOGY, John

10   Wiley & Sons, New York, NY, 1993.)

A nucleic acid of the invention can be amplified using cDNA, mRNA or alternatively, genomic DNA, as a template with appropriate oligonucleotide primers according to standard PCR amplification techniques. The nucleic acid so amplified can be cloned into an appropriate vector and characterized by DNA sequence analysis.

15   Furthermore, oligonucleotides corresponding to NOVX nucleotide sequences can be prepared by standard synthetic techniques, *e.g.*, using an automated DNA synthesizer.

As used herein, the term "oligonucleotide" refers to a series of linked nucleotide residues. A short oligonucleotide sequence may be based on, or designed from, a genomic or cDNA sequence and is used to amplify, confirm, or reveal the presence of an identical,

20   similar or complementary DNA or RNA in a particular cell or tissue. Oligonucleotides comprise a nucleic acid sequence having about 10 nt, 50 nt, or 100 nt in length, preferably about 15 nt to 30 nt in length. In one embodiment of the invention, an oligonucleotide comprising a nucleic acid molecule less than 100 nt in length would further comprise at least 6 contiguous nucleotides of SEQ ID NO:$2n$-1, wherein n is an integer between 1 and

25   124, or a complement thereof. Oligonucleotides may be chemically synthesized and may also be used as probes.

In another embodiment, an isolated nucleic acid molecule of the invention comprises a nucleic acid molecule that is a complement of the nucleotide sequence shown in SEQ ID NO:$2n$-1, wherein n is an integer between 1 and 124, or a portion of this

30   nucleotide sequence (*e.g.*, a fragment that can be used as a probe or primer or a fragment encoding a biologically-active portion of a NOVX polypeptide). A nucleic acid molecule that is complementary to the nucleotide sequence of SEQ ID NO:$2n$-1, wherein n is an integer between 1 and 124, is one that is sufficiently complementary to the nucleotide

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sequence of SEQ ID NO:2n-1, wherein n is an integer between 1 and 124, that it can hydrogen bond with few or no mismatches to the nucleotide sequence shown in SEQ ID NO:2n-1, wherein n is an integer between 1 and 124, thereby forming a stable duplex.

5   As used herein, the term "complementary" refers to Watson-Crick or Hoogsteen base pairing between nucleotides units of a nucleic acid molecule, and the term "binding" means the physical or chemical interaction between two polypeptides or compounds or associated polypeptides or compounds or combinations thereof. Binding includes ionic, non-ionic, van der Waals, hydrophobic interactions, and the like. A physical interaction can be either direct or indirect. Indirect interactions may be through or due to the effects of

10   another polypeptide or compound. Direct binding refers to interactions that do not take place through, or due to, the effect of another polypeptide or compound, but instead are without other substantial chemical intermediates.

A "fragment" provided herein is defined as a sequence of at least 6 (contiguous) nucleic acids or at least 4 (contiguous) amino acids, a length sufficient to allow for specific

15   hybridization in the case of nucleic acids or for specific recognition of an epitope in the case of amino acids, and is at most some portion less than a full length sequence. Fragments may be derived from any contiguous portion of a nucleic acid or amino acid sequence of choice.

A full-length NOVX clone is identified as containing an ATG translation start

20   codon and an in-frame stop codon. Any disclosed NOVX nucleotide sequence lacking an ATG start codon therefore encodes a truncated C-terminal fragment of the respective NOVX polypeptide, and requires that the corresponding full-length cDNA extend in the 5' direction of the disclosed sequence. Any disclosed NOVX nucleotide sequence lacking an in-frame stop codon similarly encodes a truncated N-terminal fragment of the respective

25   NOVX polypeptide, and requires that the corresponding full-length cDNA extend in the 3' direction of the disclosed sequence.

A "derivative" is a nucleic acid sequence or amino acid sequence formed from the native compounds either directly, by modification or partial substitution. An "analog" is a nucleic acid sequence or amino acid sequence that has a structure similar to, but not

30   identical to, the native compound, e.g. they differs from it in respect to certain components or side chains. Analogs may be synthetic or derived from a different evolutionary origin and may have a similar or opposite metabolic activity compared to wild type. A

"homolog" is a nucleic acid sequence or amino acid sequence of a particular gene that is derived from different species.

Derivatives and analogs may be full length or other than full length. Derivatives or analogs of the nucleic acids or proteins of the invention include, but are not limited to,

5   molecules comprising regions that are substantially homologous to the nucleic acids or proteins of the invention, in various embodiments, by at least about 70%, 80%, or 95% identity (with a preferred identity of 80-95%) over a nucleic acid or amino acid sequence of identical size or when compared to an aligned sequence in which the alignment is done by a computer homology program known in the art, or whose encoding nucleic acid is capable

10   of hybridizing to the complement of a sequence encoding the proteins under stringent, moderately stringent, or low stringent conditions. *See e.g.* Ausubel, *et al.*, CURRENT PROTOCOLS IN MOLECULAR BIOLOGY, John Wiley & Sons, New York, NY, 1993, and below.

A "homologous nucleic acid sequence" or "homologous amino acid sequence," or

15   variations thereof, refer to sequences characterized by a homology at the nucleotide level or amino acid level as discussed above. Homologous nucleotide sequences include those sequences coding for isoforms of NOVX polypeptides. Isoforms can be expressed in different tissues of the same organism as a result of, for example, alternative splicing of RNA. Alternatively, isoforms can be encoded by different genes. In the invention,

20   homologous nucleotide sequences include nucleotide sequences encoding for a NOVX polypeptide of species other than humans, including, but not limited to: vertebrates, and thus can include, *e.g.*, frog, mouse, rat, rabbit, dog, cat cow, horse, and other organisms. Homologous nucleotide sequences also include, but are not limited to, naturally occurring allelic variations and mutations of the nucleotide sequences set forth herein. A homologous

25   nucleotide sequence does not, however, include the exact nucleotide sequence encoding human NOVX protein. Homologous nucleic acid sequences include those nucleic acid sequences that encode conservative amino acid substitutions (see below) in SEQ ID NO:2n-1, wherein *n* is an integer between 1 and 124, as well as a polypeptide possessing NOVX biological activity. Various biological activities of the NOVX proteins are

30   described below.

A NOVX polypeptide is encoded by the open reading frame ("ORF") of a NOVX nucleic acid. An ORF corresponds to a nucleotide sequence that could potentially be translated into a polypeptide. A stretch of nucleic acids comprising an ORF is

uninterrupted by a stop codon. An ORF that represents the coding sequence for a full
protein begins with an ATG "start" codon and terminates with one of the three "stop"
codons, namely, TAA, TAG, or TGA. For the purposes of this invention, an ORF may be
any part of a coding sequence, with or without a start codon, a stop codon, or both. For an
5   ORF to be considered as a good candidate for coding for a *bona fide* cellular protein, a
minimum size requirement is often set, *e.g.*, a stretch of DNA that would encode a protein
of 50 amino acids or more.

The nucleotide sequences determined from the cloning of the human NOVX genes
allows for the generation of probes and primers designed for use in identifying and/or
10   cloning NOVX homologues in other cell types, *e.g.* from other tissues, as well as NOVX
homologues from other vertebrates. The probe/primer typically comprises substantially
purified oligonucleotide. The oligonucleotide typically comprises a region of nucleotide
sequence that hybridizes under stringent conditions to at least about 12, 25, 50, 100, 150,
200, 250, 300, 350 or 400 consecutive sense strand nucleotide sequence of SEQ ID
15   NO:2n-1, wherein n is an integer between 1 and 124; or an anti-sense strand nucleotide
sequence of SEQ ID NO:2n-1, wherein n is an integer between 1 and 124; or of a naturally
occurring mutant of SEQ ID NO:2n-1, wherein n is an integer between 1 and 124.

Probes based on the human NOVX nucleotide sequences can be used to detect
transcripts or genomic sequences encoding the same or homologous proteins. In various
20   embodiments, the probe has a detectable label attached, *e.g.* the label can be a radioisotope,
a fluorescent compound, an enzyme, or an enzyme co-factor. Such probes can be used as a
part of a diagnostic test kit for identifying cells or tissues which mis-express a NOVX
protein, such as by measuring a level of a NOVX-encoding nucleic acid in a sample of cells
from a subject *e.g.*, detecting NOVX mRNA levels or determining whether a genomic
25   NOVX gene has been mutated or deleted.

"A polypeptide having a biologically-active portion of a NOVX polypeptide" refers
to polypeptides exhibiting activity similar, but not necessarily identical to, an activity of a
polypeptide of the invention, including mature forms, as measured in a particular biological
assay, with or without dose dependency. A nucleic acid fragment encoding a
30   "biologically-active portion of NOVX" can be prepared by isolating a portion of SEQ ID
NO:2n-1, wherein n is an integer between 1 and 124, that encodes a polypeptide having a
NOVX biological activity (the biological activities of the NOVX proteins are described

below), expressing the encoded portion of NOVX protein (e.g., by recombinant expression *in vitro*) and assessing the activity of the encoded portion of NOVX.

NOVX Nucleic Acid and Polypeptide Variants

The invention further encompasses nucleic acid molecules that differ from the nucleotide sequences of SEQ ID NO:2n-1, wherein n is an integer between 1 and 124, due to degeneracy of the genetic code and thus encode the same NOVX proteins as that encoded by the nucleotide sequences of SEQ ID NO:2n-1, wherein n is an integer between 1 and 124. In another embodiment, an isolated nucleic acid molecule of the invention has a nucleotide sequence encoding a protein having an amino acid sequence of SEQ ID NO:2n, wherein n is an integer between 1 and 124.

In addition to the human NOVX nucleotide sequences of SEQ ID NO:2n-1, wherein n is an integer between 1 and 124, it will be appreciated by those skilled in the art that DNA sequence polymorphisms that lead to changes in the amino acid sequences of the NOVX polypeptides may exist within a population (e.g., the human population). Such genetic polymorphism in the NOVX genes may exist among individuals within a population due to natural allelic variation. As used herein, the terms "gene" and "recombinant gene" refer to nucleic acid molecules comprising an open reading frame (ORF) encoding a NOVX protein, preferably a vertebrate NOVX protein. Such natural allelic variations can typically result in 1-5% variance in the nucleotide sequence of the NOVX genes. Any and all such nucleotide variations and resulting amino acid polymorphisms in the NOVX polypeptides, which are the result of natural allelic variation and that do not alter the functional activity of the NOVX polypeptides, are intended to be within the scope of the invention.

Moreover, nucleic acid molecules encoding NOVX proteins from other species, and thus that have a nucleotide sequence that differs from a human SEQ ID NO:2n-1, wherein n is an integer between 1 and 124, are intended to be within the scope of the invention. Nucleic acid molecules corresponding to natural allelic variants and homologues of the NOVX cDNAs of the invention can be isolated based on their homology to the human NOVX nucleic acids disclosed herein using the human cDNAs, or a portion thereof, as a hybridization probe according to standard hybridization techniques under stringent hybridization conditions.

Accordingly, in another embodiment, an isolated nucleic acid molecule of the invention is at least 6 nucleotides in length and hybridizes under stringent conditions to the

nucleic acid molecule comprising the nucleotide sequence of SEQ ID NO:2n-1, wherein n is an integer between 1 and 124. In another embodiment, the nucleic acid is at least 10, 25, 50, 100, 250, 500, 750, 1000, 1500, or 2000 or more nucleotides in length. In yet another embodiment, an isolated nucleic acid molecule of the invention hybridizes to the coding

5   region. As used herein, the term "hybridizes under stringent conditions" is intended to describe conditions for hybridization and washing under which nucleotide sequences at least about 65% homologous to each other typically remain hybridized to each other.

Homologs (i.e., nucleic acids encoding NOVX proteins derived from species other than human) or other related sequences (e.g., paralogs) can be obtained by low, moderate or

10   high stringency hybridization with all or a portion of the particular human sequence as a probe using methods well known in the art for nucleic acid hybridization and cloning.

As used herein, the phrase "stringent hybridization conditions" refers to conditions under which a probe, primer or oligonucleotide will hybridize to its target sequence, but to no other sequences. Stringent conditions are sequence-dependent and will be different in

15   different circumstances. Longer sequences hybridize specifically at higher temperatures than shorter sequences. Generally, stringent conditions are selected to be about 5 °C lower than the thermal melting point (Tm) for the specific sequence at a defined ionic strength and pH. The Tm is the temperature (under defined ionic strength, pH and nucleic acid concentration) at which 50% of the probes complementary to the target sequence hybridize

20   to the target sequence at equilibrium. Since the target sequences are generally present at excess, at Tm, 50% of the probes are occupied at equilibrium. Typically, stringent conditions will be those in which the salt concentration is less than about 1.0 M sodium ion, typically about 0.01 to 1.0 M sodium ion (or other salts) at pH 7.0 to 8.3 and the temperature is at least about 30 °C for short probes, primers or oligonucleotides (e.g., 10 nt

25   to 50 nt) and at least about 60 °C for longer probes, primers and oligonucleotides. Stringent conditions may also be achieved with the addition of destabilizing agents, such as formamide.

Stringent conditions are known to those skilled in the art and can be found in Ausubel, et al., (eds.), CURRENT PROTOCOLS IN MOLECULAR BIOLOGY, John Wiley & Sons,

30   N.Y. (1989), 6.3.1-6.3.6. Preferably, the conditions are such that sequences at least about 65%, 70%, 75%, 85%, 90%, 95%, 98%, or 99% homologous to each other typically remain hybridized to each other. A non-limiting example of stringent hybridization conditions are hybridization in a high salt buffer comprising 6X SSC, 50 mM Tris-HCl (pH 7.5), 1 mM

EDTA, 0.02% PVP, 0.02% Ficoll, 0.02% BSA, and 500 mg/ml denatured salmon sperm DNA at 65°C, followed by one or more washes in 0.2X SSC, 0.01% BSA at 50°C. An isolated nucleic acid molecule of the invention that hybridizes under stringent conditions to a sequence of SEQ ID NO:$2n$-1, wherein n is an integer between 1 and 124, corresponds to

5 a naturally-occurring nucleic acid molecule. As used herein, a "naturally-occurring" nucleic acid molecule refers to an RNA or DNA molecule having a nucleotide sequence that occurs in nature (e.g., encodes a natural protein).

In a second embodiment, a nucleic acid sequence that is hybridizable to the nucleic acid molecule comprising the nucleotide sequence of SEQ ID NO:$2n$-1, wherein n is an

10 integer between 1 and 124, or fragments, analogs or derivatives thereof, under conditions of moderate stringency is provided. A non-limiting example of moderate stringency hybridization conditions are hybridization in 6X SSC, 5X Reinhardt's solution, 0.5% SDS and 100 mg/ml denatured salmon sperm DNA at 55 °C, followed by one or more washes in 1X SSC, 0.1% SDS at 37 °C. Other conditions of moderate stringency that may be used

15 are well-known within the art. See, e.g., Ausubel, et al. (eds.), 1993, CURRENT PROTOCOLS IN MOLECULAR BIOLOGY, John Wiley & Sons, NY, and Krieger, 1990; GENE TRANSFER AND EXPRESSION, A LABORATORY MANUAL, Stockton Press, NY.

In a third embodiment, a nucleic acid that is hybridizable to the nucleic acid molecule comprising the nucleotide sequences of SEQ ID NO:$2n$-1, wherein n is an integer

20 between 1 and 124, or fragments, analogs or derivatives thereof, under conditions of low stringency, is provided. A non-limiting example of low stringency hybridization conditions are hybridization in 35% formamide, 5X SSC, 50 mM Tris-HCl (pH 7.5), 5 mM EDTA, 0.02% PVP, 0.02% Ficoll, 0.2% BSA, 100 mg/ml denatured salmon sperm DNA, 10% (wt/vol) dextran sulfate at 40°C, followed by one or more washes in 2X SSC, 25 mM

25 Tris-HCl (pH 7.4), 5 mM EDTA, and 0.1% SDS at 50°C. Other conditions of low stringency that may be used are well known in the art (e.g., as employed for cross-species hybridizations). See, e.g., Ausubel, et al. (eds.), 1993, CURRENT PROTOCOLS IN MOLECULAR BIOLOGY, John Wiley & Sons, NY, and Kriegler, 1990, GENE TRANSFER AND EXPRESSION, A LABORATORY MANUAL, Stockton Press, NY; Shilo and Weinberg, 1981.

30 *Proc Natl Acad Sci USA* **78**: 6789-6792.

Conservative Mutations

In addition to naturally-occurring allelic variants of NOVX sequences that may exist in the population, the skilled artisan will further appreciate that changes can be

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introduced by mutation into the nucleotide sequences of SEQ ID NO:$2n$-1, wherein n is an integer between 1 and 124, thereby leading to changes in the amino acid sequences of the encoded NOVX protein, without altering the functional ability of that NOVX protein. For example, nucleotide substitutions leading to amino acid substitutions at "non-essential"

5 amino acid residues can be made in the sequence of SEQ ID NO:$2n$, wherein n is an integer between 1 and 124. A "non-essential" amino acid residue is a residue that can be altered from the wild-type sequences of the NOVX proteins without altering their biological activity, whereas an "essential" amino acid residue is required for such biological activity. For example, amino acid residues that are conserved among the NOVX proteins of the

10 invention are predicted to be particularly non-amenable to alteration. Amino acids for which conservative substitutions can be made are well-known within the art.

Another aspect of the invention pertains to nucleic acid molecules encoding NOVX proteins that contain changes in amino acid residues that are not essential for activity. Such NOVX proteins differ in amino acid sequence from SEQ ID NO:$2n$-1, wherein n is an

15 integer between 1 and 124, yet retain biological activity. In one embodiment, the isolated nucleic acid molecule comprises a nucleotide sequence encoding a protein, wherein the protein comprises an amino acid sequence at least about 40% homologous to the amino acid sequences of SEQ ID NO:$2n$, wherein n is an integer between 1 and 124. Preferably, the protein encoded by the nucleic acid molecule is at least about 60% homologous to SEQ

20 ID NO:$2n$, wherein n is an integer between 1 and 124; more preferably at least about 70% homologous to SEQ ID NO:$2n$, wherein n is an integer between 1 and 124; still more preferably at least about 80% homologous to SEQ ID NO:$2n$, wherein n is an integer between 1 and 124; even more preferably at least about 90% homologous to SEQ ID NO:$2n$, wherein n is an integer between 1 and 124; and most preferably at least about 95%

25 homologous to SEQ ID NO:$2n$, wherein n is an integer between 1 and 124.

An isolated nucleic acid molecule encoding a NOVX protein homologous to the protein of SEQ ID NO:$2n$, wherein n is an integer between 1 and 124, can be created by introducing one or more nucleotide substitutions, additions or deletions into the nucleotide sequence of SEQ ID NO:$2n$-1, wherein n is an integer between 1 and 124, such that one or

30 more amino acid substitutions, additions or deletions are introduced into the encoded protein.

Mutations can be introduced any one of SEQ ID NO:$2n$-1, wherein n is an integer between 1 and 124, by standard techniques, such as site-directed mutagenesis and

PCR-mediated mutagenesis. Preferably, conservative amino acid substitutions are made at one or more predicted, non-essential amino acid residues. A "conservative amino acid substitution" is one in which the amino acid residue is replaced with an amino acid residue having a similar side chain. Families of amino acid residues having similar side chains

5 have been defined within the art. These families include amino acids with basic side chains (e.g., lysine, arginine, histidine), acidic side chains (e.g., aspartic acid, glutamic acid), uncharged polar side chains (e.g., glycine, asparagine, glutamine, serine, threonine, tyrosine, cysteine), nonpolar side chains (e.g., alanine, valine, leucine, isoleucine, proline, phenylalanine, methionine, tryptophan), beta-branched side chains (e.g., threonine, valine,

10 isoleucine) and aromatic side chains (e.g., tyrosine, phenylalanine, tryptophan, histidine). Thus, a predicted non-essential amino acid residue in the NOVX protein is replaced with another amino acid residue from the same side chain family. Alternatively, in another embodiment, mutations can be introduced randomly along all or part of a NOVX coding sequence, such as by saturation mutagenesis, and the resultant mutants can be screened for

15 NOVX biological activity to identify mutants that retain activity. Following mutagenesis of a nucleic acid of SEQ ID NO:$2n$-1, wherein n is an integer between 1 and 124, the encoded protein can be expressed by any recombinant technology known in the art and the activity of the protein can be determined.

The relatedness of amino acid families may also be determined based on side chain

20 interactions. Substituted amino acids may be fully conserved "strong" residues or fully conserved "weak" residues. The "strong" group of conserved amino acid residues may be any one of the following groups: STA, NEQK, NHQK, NDEQ, QHRK, MILV, MILF, HY, FYW, wherein the single letter amino acid codes are grouped by those amino acids that may be substituted for each other. Likewise, the "weak" group of conserved residues may

25 be any one of the following: CSA, ATV, SAG, STNK, STPA, SGND, SNDEQK, NDEQHK, NEQHRK, HFY, wherein the letters within each group represent the single letter amino acid code.

In one embodiment, a mutant NOVX protein can be assayed for (i) the ability to form protein:protein interactions with other NOVX proteins, other cell-surface proteins, or

30 biologically-active portions thereof, (ii) complex formation between a mutant NOVX protein and a NOVX ligand; or (iii) the ability of a mutant NOVX protein to bind to an intracellular target protein or biologically-active portion thereof; (e.g. avidin proteins).

In yet another embodiment, a mutant NOVX protein can be assayed for the ability to regulate a specific biological function (*e.g.*, regulation of insulin release).

Interfering RNA

In one aspect of the invention, NOVX gene expression can be attenuated by RNA
5   interference. One approach well-known in the art is short interfering RNA (siRNA)
mediated gene silencing where expression products of a NOVX gene are targeted by
specific double stranded NOVX derived siRNA nucleotide sequences that are
complementary to at least a 19-25 nt long segment of the NOVX gene transcript, including
the 5' untranslated (UT) region, the ORF, or the 3' UT region. *See, e.g.*, PCT applications
10   WO00/44895, WO99/32619, WO01/75164, WO01/92513, WO 01/29058, WO01/89304,
WO02/16620, and WO02/29858, each incorporated by reference herein in their entirety.
Targeted genes can be a NOVX gene, or an upstream or downstream modulator of the
NOVX gene. Nonlimiting examples of upstream or downstream modulators of a NOVX
gene include, *e.g.*, a transcription factor that binds the NOVX gene promoter, a kinase or
15   phosphatase that interacts with a NOVX polypeptide, and polypeptides involved in a
NOVX regulatory pathway.

According to the methods of the present invention, NOVX gene expression is
silenced using short interfering RNA. A NOVX polynucleotide according to the invention
includes a siRNA polynucleotide. Such a NOVX siRNA can be obtained using a NOVX
20   polynucleotide sequence, for example, by processing the NOVX ribopolynucleotide
sequence in a cell-free system, such as but not limited to a Drosophila extract, or by
transcription of recombinant double stranded NOVX RNA or by chemical synthesis of
nucleotide sequences homologous to a NOVX sequence. *See, e.g.*, Tuschl, Zamore,
Lehmann, Bartel and Sharp (1999), Genes & Dev. 13: 3191-3197, incorporated herein by
25   reference in its entirety. When synthesized, a typical 0.2 micromolar-scale RNA synthesis
provides about 1 milligram of siRNA, which is sufficient for 1000 transfection experiments
using a 24-well tissue culture plate format.

The most efficient silencing is generally observed with siRNA duplexes composed
of a 21-nt sense strand and a 21-nt antisense strand, paired in a manner to have a 2-nt
30   3' overhang. The sequence of the 2-nt 3' overhang makes an additional small contribution
to the specificity of siRNA target recognition. The contribution to specificity is localized to
the unpaired nucleotide adjacent to the first paired bases. In one embodiment, the
nucleotides in the 3' overhang are ribonucleotides. In an alternative embodiment, the

nucleotides in the 3' overhang are deoxyribonucleotides. "Using 2'-deoxyribonucleotides in the 3' overhangs is as efficient as using ribonucleotides, but deoxyribonucleotides are often cheaper to synthesize and are most likely more nuclease resistant.

A contemplated recombinant expression vector of the invention comprises a NOVX
5   DNA molecule cloned into an expression vector comprising operatively-linked regulatory sequences flanking the NOVX sequence in a manner that allows for expression (by transcription of the DNA molecule) of both strands. An RNA molecule that is antisense to NOVX mRNA is transcribed by a first promoter (e.g., a promoter sequence 3' of the cloned DNA) and an RNA molecule that is the sense strand for the NOVX mRNA is transcribed
10   by a second promoter (e.g., a promoter sequence 5' of the cloned DNA). The sense and antisense strands may hybridize in vivo to generate siRNA constructs for silencing of the NOVX gene. Alternatively, two constructs can be utilized to create the sense and anti-sense strands of a siRNA construct. Finally, cloned DNA can encode a construct having secondary structure, wherein a single transcript has both the sense and
15   complementary antisense sequences from the target gene or genes. In an example of this embodiment, a hairpin RNAi product is homologous to all or a portion of the target gene. In another example, a hairpin RNAi product is a siRNA. The regulatory sequences flanking the NOVX sequence may be identical or may be different, such that their expression may be modulated independently, or in a temporal or spatial manner.

20   In a specific embodiment, siRNAs are transcribed intracellularly by cloning the NOVX gene templates into a vector containing, e.g., a RNA pol III transcription unit from the smaller nuclear RNA (snRNA) U6 or the human RNase P RNA H1. One example of a vector system is the GeneSuppressor™ RNA Interference kit (commercially available from Imgenex). The U6 and H1 promoters are members of the type III class of Pol III promoters.
25   The +1 nucleotide of the U6-like promoters is always guanosine, whereas the +1 for H1 promoters is adenosine. The termination signal for these promoters is defined by five consecutive thymidines. The transcript is typically cleaved after the second uridine. Cleavage at this position generates a 3' UU overhang in the expressed siRNA, which is similar to the 3' overhangs of synthetic siRNAs. Any sequence less than 400 nucleotides in
30   length can be transcribed by these promoter, therefore they are ideally suited for the expression of around 21-nucleotide siRNAs in, e.g., an approximately 50-nucleotide RNA stem-loop transcript.

A siRNA vector appears to have an advantage over synthetic siRNAs where long-term knock-down of expression is desired. Cells transfected with a siRNA expression vector would experience steady, long-term mRNA inhibition. In contrast, cells transfected with exogenous synthetic siRNAs typically recover from mRNA suppression within seven days or ten rounds of cell division. The long-term gene silencing ability of siRNA expression vectors may provide for applications in gene therapy.

In general, siRNAs are chopped from longer dsRNA by an ATP-dependent ribonuclease called DICER. DICER is a member of the RNase III family of double-stranded RNA-specific endonucleases. The siRNAs assemble with cellular proteins into an endonuclease complex. *In vitro* studies in Drosophila suggest that the siRNAs/protein complex (siRNP) is then transferred to a second enzyme complex, called an RNA-induced silencing complex (RISC), which contains an endoribonuclease that is distinct from DICER. RISC uses the sequence encoded by the antisense siRNA strand to find and destroy mRNAs of complementary sequence. The siRNA thus acts as a guide, restricting the ribonuclease to cleave only mRNAs complementary to one of the two siRNA strands.

A NOVX mRNA region to be targeted by siRNA is generally selected from a desired NOVX sequence beginning 50 to100 nt downstream of the start codon. Alternatively, 5' or 3' UTRs and regions nearby the start codon can be used but are generally avoided, as these may be richer in regulatory protein binding sites. UTR-binding proteins and/or translation initiation complexes may interfere with binding of the siRNP or RISC endonuclease complex. An initial BLAST homology search for the selected siRNA sequence is done against an available nucleotide sequence library to ensure that only one gene is targeted. Specificity of target recognition by siRNA duplexes indicate that a single point mutation located in the paired region of an siRNA duplex is sufficient to abolish target mRNA degradation. See, Elbashir *et al.* 2001 EMBO J. 20(23):6877-88. Hence, consideration should be taken to accommodate SNPs, polymorphisms, allelic variants or species-specific variations when targeting a desired gene.

In one embodiment, a complete NOVX siRNA experiment includes the proper negative control. A negative control siRNA generally has the same nucleotide composition as the NOVX siRNA but lack significant sequence homology to the genome. Typically, one would scramble the nucleotide sequence of the NOVX siRNA and do a homology search to make sure it lacks homology to any other gene.

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Two independent NOVX siRNA duplexes can be used to knock-down a target NOVX gene. This helps to control for specificity of the silencing effect. In addition, expression of two independent genes can be simultaneously knocked down by using equal concentrations of different NOVX siRNA duplexes, *e.g.*, a NOVX siRNA and an siRNA

5   for a regulator of a NOVX gene or polypeptide. Availability of siRNA-associating proteins is believed to be more limiting than target mRNA accessibility.

A targeted NOVX region is typically a sequence of two adenines (AA) and two thymidines (TT) divided by a spacer region of nineteen (N19) residues (*e.g.*, AA(N19)TT). A desirable spacer region has a G/C-content of approximately 30% to 70%, and more

10   preferably of about 50%. If the sequence AA(N19)TT is not present in the target sequence, an alternative target region would be AA(N21). The sequence of the NOVX sense siRNA corresponds to (N19)TT or N21, respectively. In the latter case, conversion of the 3' end of the sense siRNA to TT can be performed if such a sequence does not naturally occur in the NOVX polynucleotide. The rationale for this sequence conversion is to generate a

15   symmetric duplex with respect to the sequence composition of the sense and antisense 3' overhangs. Symmetric 3' overhangs may help to ensure that the siRNPs are formed with approximately equal ratios of sense and antisense target RNA-cleaving siRNPs. *See, e.g.*, Elbashir, Lendeckel and Tuschl (2001). Genes & Dev. 15: 188-200, incorporated by reference herein in its entirely. The modification of the overhang of the sense sequence of

20   the siRNA duplex is not expected to affect targeted mRNA recognition, as the antisense siRNA strand guides target recognition.

Alternatively, if the NOVX target mRNA does not contain a suitable AA(N21) sequence, one may search for the sequence NA(N21). Further, the sequence of the sense strand and antisense strand may still be synthesized as 5' (N19)TT, as it is believed that the

25   sequence of the 3'-most nucleotide of the antisense siRNA does not contribute to specificity. Unlike antisense or ribozyme technology, the secondary structure of the target mRNA does not appear to have a strong effect on silencing. *See,* Harborth, *et al.* (2001) J. Cell Science 114: 4557-4565, incorporated by reference in its entirety.

Transfection of NOVX siRNA duplexes can be achieved using standard nucleic

30   acid transfection methods, for example, OLIGOFECTAMINE Reagent (commercially available from Invitrogen). An assay for NOVX gene silencing is generally performed approximately 2 days after transfection. No NOVX gene silencing has been observed in the absence of transfection reagent, allowing for a comparative analysis of the wild-type

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and silenced NOVX phenotypes. In a specific embodiment, for one well of a 24-well plate, approximately 0.84 μg of the siRNA duplex is generally sufficient. Cells are typically seeded the previous day, and are transfected at about 50% confluence. The choice of cell culture media and conditions are routine to those of skill in the art, and will vary with the

5   choice of cell type. The efficiency of transfection may depend on the cell type, but also on the passage number and the confluency of the cells. The time and the manner of formation of siRNA-liposome complexes (e.g. inversion versus vortexing) are also critical. Low transfection efficiencies are the most frequent cause of unsuccessful NOVX silencing. The efficiency of transfection needs to be carefully examined for each new cell line to be used.

10   Preferred cell are derived from a mammal, more preferably from a rodent such as a rat or mouse, and most preferably from a human. Where used for therapeutic treatment, the cells are preferentially autologous, although non-autologous cell sources are also contemplated as within the scope of the present invention.

For a control experiment, transfection of 0.84 μg single-stranded sense NOVX

15   siRNA will have no effect on NOVX silencing, and 0.84 μg antisense siRNA has a weak silencing effect when compared to 0.84 μg of duplex siRNAs. Control experiments again allow for a comparative analysis of the wild-type and silenced NOVX phenotypes. To control for transfection efficiency, targeting of common proteins is typically performed, for example targeting of lamin A/C or transfection of a CMV-driven EGFP-expression plasmid

20   (e.g. commercially available from Clontech). In the above example, a determination of the fraction of lamin A/C knockdown in cells is determined the next day by such techniques as immunofluorescence, Western blot, Northern blot or other similar assays for protein expression or gene expression. Lamin A/C monoclonal antibodies may be obtained from Santa Cruz Biotechnology.

25   Depending on the abundance and the half life (or turnover) of the targeted NOVX polynucleotide in a cell, a knock-down phenotype may become apparent after 1 to 3 days, or even later. In cases where no NOVX knock-down phenotype is observed, depletion of the NOVX polynucleotide may be observed by immunofluorescence or Western blotting. If the NOVX polynucleotide is still abundant after 3 days, cells need to be split and

30   transferred to a fresh 24-well plate for re-transfection. If no knock-down of the targeted protein is observed, it may be desirable to analyze whether the target mRNA (NOVX or a NOVX upstream or downstream gene) was effectively destroyed by the transfected siRNA duplex. Two days after transfection, total RNA is prepared, reverse transcribed using a

target-specific primer, and PCR-amplified with a primer pair covering at least one exon-exon junction in order to control for amplification of pre-mRNAs. RT/PCR of a non-targeted mRNA is also needed as control. Effective depletion of the mRNA yet undetectable reduction of target protein may indicate that a large reservoir of stable NOVX

5   protein may exist in the cell. Multiple transfection in sufficiently long intervals may be necessary until the target protein is finally depleted to a point where a phenotype may become apparent. If multiple transfection steps are required, cells are split 2 to 3 days after transfection. The cells may be transfected immediately after splitting.

An inventive therapeutic method of the invention contemplates administering a

10   NOVX siRNA construct as therapy to compensate for increased or aberrant NOVX expression or activity. The NOVX ribopolynucleotide is obtained and processed into siRNA fragments, or a NOVX siRNA is synthesized, as described above. The NOVX siRNA is administered to cells or tissues using known nucleic acid transfection techniques, as described above. A NOVX siRNA specific for a NOVX gene will decrease or

15   knockdown NOVX transcription products, which will lead to reduced NOVX polypeptide production, resulting in reduced NOVX polypeptide activity in the cells or tissues.

The present invention also encompasses a method of treating a disease or condition associated with the presence of a NOVX protein in an individual comprising administering to the individual an RNAi construct that targets the mRNA of the protein (the mRNA that

20   encodes the protein) for degradation. A specific RNAi construct includes a siRNA or a double stranded gene transcript that is processed into siRNAs. Upon treatment, the target protein is not produced or is not produced to the extent it would be in the absence of the treatment.

Where the NOVX gene function is not correlated with a known phenotype, a

25   control sample of cells or tissues from healthy individuals provides a reference standard for determining NOVX expression levels. Expression levels are detected using the assays described, e.g., RT-PCR, Northern blotting, Western blotting, ELISA, and the like. A subject sample of cells or tissues is taken from a mammal, preferably a human subject, suffering from a disease state. The NOVX ribopolynucleotide is used to produce siRNA

30   constructs, that are specific for the NOVX gene product. These cells or tissues are treated by administering NOVX siRNA's to the cells or tissues by methods described for the transfection of nucleic acids into a cell or tissue, and a change in NOVX polypeptide or polynucleotide expression is observed in the subject sample relative to the control sample,

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using the assays described. This NOVX gene knockdown approach provides a rapid method for determination of a NOVX minus (NOVX⁻) phenotype in the treated subject sample. The NOVX⁻ phenotype observed in the treated subject sample thus serves as a marker for monitoring the course of a disease state during treatment.

5   In specific embodiments, a NOVX siRNA is used in therapy. Methods for the generation and use of a NOVX siRNA are known to those skilled in the art. Example techniques are provided below.

Production of RNAs

          Sense RNA (ssRNA) and antisense RNA (asRNA) of NOVX are produced using
10   known methods such as transcription in RNA expression vectors. In the initial experiments, the sense and antisense RNA are about 500 bases in length each. The produced ssRNA and asRNA (0.5 µM) in 10 mM Tris-HCl (pH 7.5) with 20 mM NaCl were heated to 95° C for 1 min then cooled and annealed at room temperature for 12 to 16 h. The RNAs are precipitated and resuspended in lysis buffer (below). To monitor
15   annealing, RNAs are electrophoresed in a 2% agarose gel in TBE buffer and stained with ethidium bromide. See, e.g., Sambrook et al., Molecular Cloning. Cold Spring Harbor Laboratory Press, Plainview, N.Y. (1989).

Lysate Preparation

          Untreated rabbit reticulocyte lysate (Ambion) are assembled according to the
20   manufacturer's directions. dsRNA is incubated in the lysate at 30° C for 10 min prior to the addition of mRNAs. Then NOVX mRNAs are added and the incubation continued for an additional 60 min. The molar ratio of double stranded RNA and mRNA is about 200:1. The NOVX mRNA is radiolabeled (using known techniques) and its stability is monitored by gel electrophoresis.

25   In a parallel experiment made with the same conditions, the double stranded RNA is internally radiolabeled with a ³²P-ATP. Reactions are stopped by the addition of 2 X proteinase K buffer and deproteinized as described previously (Tuschl et al., Genes Dev., 13:3191-3197 (1999)). Products are analyzed by electrophoresis in 15% or 18% polyacrylamide sequencing gels using appropriate RNA standards. By monitoring the gels
30   for radioactivity, the natural production of 10 to 25 nt RNAs from the double stranded RNA can be determined.

The band of double stranded RNA, about 21-23 bps, is eluded. The efficacy of these 21-23 mers for suppressing NOVX transcription is assayed in vitro using the same rabbit reticulocyte assay described above using 50 nanomolar of double stranded 21-23 mer for each assay. The sequence of these 21-23 mers is then determined using standard

5   nucleic acid sequencing techniques.

RNA Preparation

21 nt RNAs, based on the sequence determined above, are chemically synthesized using Expedite RNA phosphoramidites and thymidine phosphoramidite (Proligo, Germany). Synthetic oligonucleotides are deprotected and gel-purified (Elbashir,

10   Lendeckel, & Tuschl, Genes & Dev. 15, 188-200 (2001)), followed by Sep-Pak C18 cartridge (Waters, Milford, Mass., USA) purification (Tuschl, et al., Biochemistry, 32:11658-11668 (1993)).

These RNAs (20 μM) single strands are incubated in annealing buffer (100 mM potassium acetate, 30 mM HEPES-KOH at pH 7.4, 2 mM magnesium acetate) for 1 min at

15   90° C followed by 1 h at 37° C.

Cell Culture

A cell culture known in the art to regularly express NOVX is propagated using standard conditions. 24 hours before transfection, at approx. 80% confluency, the cells are trypsinized and diluted 1:5 with fresh medium without antibiotics (1-3 X 105 cells/ml) and

20   transferred to 24-well plates (500 ml/well). Transfection is performed using a commercially available lipofection kit and NOVX expression is monitored using standard techniques with positive and negative control. A positive control is cells that naturally express NOVX while a negative control is cells that do not express NOVX. Base-paired 21 and 22 nt siRNAs with overhanging 3' ends mediate efficient sequence-specific mRNA

25   degradation in lysates and in cell culture. Different concentrations of siRNAs are used. An efficient concentration for suppression in vitro in mammalian culture is between 25 nM to 100 nM final concentration. This indicates that siRNAs are effective at concentrations that are several orders of magnitude below the concentrations applied in conventional antisense or ribozyme gene targeting experiments.

30   The above method provides a way both for the deduction of NOVX siRNA sequence and the use of such siRNA for in vitro suppression. In vivo suppression may be

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performed using the same siRNA using well known in vivo transfection or gene therapy transfection techniques.

Antisense Nucleic Acids

Another aspect of the invention pertains to isolated antisense nucleic acid molecules

5   that are hybridizable to or complementary to the nucleic acid molecule comprising the nucleotide sequence of SEQ ID NO:$2n$-1, wherein n is an integer between 1 and 124, or fragments, analogs or derivatives thereof. An "antisense" nucleic acid comprises a nucleotide sequence that is complementary to a "sense" nucleic acid encoding a protein (*e.g.*, complementary to the coding strand of a double-stranded cDNA molecule or

10   complementary to an mRNA sequence). In specific aspects, antisense nucleic acid molecules are provided that comprise a sequence complementary to at least about 10, 25, 50, 100, 250 or 500 nucleotides or an entire NOVX coding strand, or to only a portion thereof. Nucleic acid molecules encoding fragments, homologs, derivatives and analogs of a NOVX protein of SEQ ID NO:$2n$, wherein n is an integer between 1 and 124, or

15   antisense nucleic acids complementary to a NOVX nucleic acid sequence of SEQ ID NO:$2n$-1, wherein n is an integer between 1 and 124, are additionally provided.

In one embodiment, an antisense nucleic acid molecule is antisense to a "coding region" of the coding strand of a nucleotide sequence encoding a NOVX protein. The term "coding region" refers to the region of the nucleotide sequence comprising codons which

20   are translated into amino acid residues. In another embodiment, the antisense nucleic acid molecule is antisense to a "noncoding region" of the coding strand of a nucleotide sequence encoding the NOVX protein. The term "noncoding region" refers to 5' and 3' sequences which flank the coding region that are not translated into amino acids (*i.e.*, also referred to as 5' and 3' untranslated regions).

25   Given the coding strand sequences encoding the NOVX protein disclosed herein, antisense nucleic acids of the invention can be designed according to the rules of Watson and Crick or Hoogsteen base pairing. The antisense nucleic acid molecule can be complementary to the entire coding region of NOVX mRNA, but more preferably is an oligonucleotide that is antisense to only a portion of the coding or noncoding region of

30   NOVX mRNA. For example, the antisense oligonucleotide can be complementary to the region surrounding the translation start site of NOVX mRNA. An antisense oligonucleotide can be, for example, about 5, 10, 15, 20, 25, 30, 35, 40, 45 or 50 nucleotides in length. An antisense nucleic acid of the invention can be constructed using

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chemical synthesis or enzymatic ligation reactions using procedures known in the art. For example, an antisense nucleic acid (*e.g.*, an antisense oligonucleotide) can be chemically synthesized using naturally-occurring nucleotides or variously modified nucleotides designed to increase the biological stability of the molecules or to increase the physical

5   stability of the duplex formed between the antisense and sense nucleic acids (*e.g.*, phosphorothioate derivatives and acridine substituted nucleotides can be used).

Examples of modified nucleotides that can be used to generate the antisense nucleic acid include: 5-fluorouracil, 5-bromouracil, 5-chlorouracil, 5-iodouracil, hypoxanthine, xanthine, 4-acetylcytosine, 5-carboxymethylaminomethyl-2-thiouridine,

10   5-(carboxyhydroxylmethyl) uracil, 5-carboxymethylaminomethyluracil, dihydrouracil, beta-D-galactosylqueosine, inosine, N6-isopentenyladenine, 1-methylguanine, 1-methylinosine, 2,2-dimethylguanine, 2-methyladenine, 2-methylguanine, 5-methoxyuracil, 3-methylcytosine, 5-methylcytosine, N6-adenine, 7-methylguanine, 5-methylaminomethyluracil, 5-methoxyaminomethyl-2-thiouracil, 2-thiouracil,

15   4-thiouracil, beta-D-mannosylqueosine, 5'-methoxycarboxymethyluracil, 2-methylthio-N6-isopentenyladenine, uracil-5-oxyacetic acid (v), wybutoxosine, pseudouracil, queosine, 2-thiocytosine, 5-methyl-2-thiouracil, 5-methyluracil, uracil-5-oxyacetic acid methylester, uracil-5-oxyacetic acid (v), 5-methyl-2-thiouracil, 3-(3-amino-3-N-2-carboxypropyl) uracil, (acp3)w, and 2,6-diaminopurine. Alternatively,

20   the antisense nucleic acid can be produced biologically using an expression vector into which a nucleic acid has been subcloned in an antisense orientation (*i.e.*, RNA transcribed from the inserted nucleic acid will be of an antisense orientation to a target nucleic acid of interest, described further in the following subsection).

The antisense nucleic acid molecules of the invention are typically administered to a

25   subject or generated *in situ* such that they hybridize with or bind to cellular mRNA and/or genomic DNA encoding a NOVX protein to thereby inhibit expression of the protein (*e.g.*, by inhibiting transcription and/or translation). The hybridization can be by conventional nucleotide complementarity to form a stable duplex, or, for example, in the case of an antisense nucleic acid molecule that binds to DNA duplexes, through specific interactions

30   in the major groove of the double helix. An example of a route of administration of antisense nucleic acid molecules of the invention includes direct injection at a tissue site. Alternatively, antisense nucleic acid molecules can be modified to target selected cells and then administered systemically. For example, for systemic administration, antisense

molecules can be modified such that they specifically bind to receptors or antigens expressed on a selected cell surface (*e.g.*, by linking the antisense nucleic acid molecules to peptides or antibodies that bind to cell surface receptors or antigens). The antisense nucleic acid molecules can also be delivered to cells using the vectors described herein. To achieve

5   sufficient nucleic acid molecules, vector constructs in which the antisense nucleic acid molecule is placed under the control of a strong pol II or pol III promoter are preferred.

In yet another embodiment, the antisense nucleic acid molecule of the invention is an α-anomeric nucleic acid molecule. An α-anomeric nucleic acid molecule forms specific double-stranded hybrids with complementary RNA in which, contrary to the usual β-units,

10   the strands run parallel to each other. *See, e.g.,* Gaultier, *et al.,* 1987. *Nucl. Acids Res.* **15**: 6625-6641. The antisense nucleic acid molecule can also comprise a 2'-o-methylribonucleotide (*See, e.g.,* Inoue, *et al.* 1987. *Nucl. Acids Res.* **15**: 6131-6148) or a chimeric RNA-DNA analogue (*See, e.g.,* Inoue, *et al.,* 1987. *FEBS Lett.* **215**: 327-330.

Ribozymes and PNA Moieties

15   Nucleic acid modifications include, by way of non-limiting example, modified bases, and nucleic acids whose sugar phosphate backbones are modified or derivatized. These modifications are carried out at least in part to enhance the chemical stability of the modified nucleic acid, such that they may be used, for example, as antisense binding nucleic acids in therapeutic applications in a subject.

20   In one embodiment, an antisense nucleic acid of the invention is a ribozyme. Ribozymes are catalytic RNA molecules with ribonuclease activity that are capable of cleaving a single-stranded nucleic acid, such as an mRNA, to which they have a complementary region. Thus, ribozymes (*e.g.*, hammerhead ribozymes as described in Haselhoff and Gerlach 1988. *Nature* **334**: 585-591) can be used to catalytically cleave

25   NOVX mRNA transcripts to thereby inhibit translation of NOVX mRNA. A ribozyme having specificity for a NOVX-encoding nucleic acid can be designed based upon the nucleotide sequence of a NOVX cDNA disclosed herein (*i.e.,* SEQ ID NO:$2n$-1, wherein n is an integer between 1 and 124). For example, a derivative of a *Tetrahymena* L-19 IVS RNA can be constructed in which the nucleotide sequence of the active site is

30   complementary to the nucleotide sequence to be cleaved in a NOVX-encoding mRNA. *See, e.g.,* U.S. Patent 4,987,071 to Cech, *et al.* and U.S. Patent 5,116,742 to Cech, *et al.* NOVX mRNA can also be used to select a catalytic RNA having a specific ribonuclease

activity from a pool of RNA molecules. *See, e.g.,* Bartel *et al.*, (1993) *Science* 261:1411-1418.

Alternatively, NOVX gene expression can be inhibited by targeting nucleotide sequences complementary to the regulatory region of the NOVX nucleic acid (*e.g.,* the

5   NOVX promoter and/or enhancers) to form triple helical structures that prevent transcription of the NOVX gene in target cells. *See, e.g.,* Helene, 1991. *Anticancer Drug Des.* 6: 569-84; Helene, *et al.* 1992. *Ann. N.Y. Acad. Sci.* 660: 27-36; Maher, 1992. *Bioassays* 14: 807-15.

. In various embodiments, the NOVX nucleic acids can be modified at the base

10   moiety, sugar moiety or phosphate backbone to improve, *e.g.,* the stability, hybridization, or solubility of the molecule. For example, the deoxyribose phosphate backbone of the nucleic acids can be modified to generate peptide nucleic acids. *See, e.g.,* Hyrup, *et al.,* 1996. *Bioorg Med Chem* 4: 5-23. As used herein, the terms "peptide nucleic acids" or "PNAs" refer to nucleic acid mimics (*e.g.,* DNA mimics) in which the deoxyribose

15   phosphate backbone is replaced by a pseudopeptide backbone and only the four natural nucleotide bases are retained. The neutral backbone of PNAs has been shown to allow for specific hybridization to DNA and RNA under conditions of low ionic strength. The synthesis of PNA oligomer can be performed using standard solid phase peptide synthesis protocols as described in Hyrup, *et al.,* 1996. *supra*; Perry-O'Keefe, *et al.,* 1996. *Proc. Natl.*

20   *Acad. Sci. USA* 93: 14670-14675.

PNAs of NOVX can be used in therapeutic and diagnostic applications. For example, PNAs can be used as antisense or antigene agents for sequence-specific modulation of gene expression by, *e.g.,* inducing transcription or translation arrest or inhibiting replication. PNAs of NOVX can also be used, for example, in the analysis of

25   single base pair mutations in a gene (*e.g.,* PNA directed PCR clamping; as artificial restriction enzymes when used in combination with other enzymes, *e.g.,* S_1 nucleases (*See,* Hyrup, *et al.,* 1996.*supra*); or as probes or primers for DNA sequence and hybridization (*See,* Hyrup, *et al.,* 1996, *supra*; Perry-O'Keefe, *et al.,* 1996. *supra*).

In another embodiment, PNAs of NOVX can be modified, *e.g.,* to enhance their

30   stability or cellular uptake, by attaching lipophilic or other helper groups to PNA, by the formation of PNA-DNA chimeras, or by the use of liposomes or other techniques of drug delivery known in the art. For example, PNA-DNA chimeras of NOVX can be generated that may combine the advantageous properties of PNA and DNA. Such chimeras allow

DNA recognition enzymes (*e.g.*, RNase H and DNA polymerases) to interact with the DNA portion while the PNA portion would provide high binding affinity and specificity. PNA-DNA chimeras can be linked using linkers of appropriate lengths selected in terms of base stacking, number of bonds between the nucleotide bases, and orientation (*see*, Hyrup,

5   et al., 1996. *supra*). The synthesis of PNA-DNA chimeras can be performed as described in Hyrup, *et al.*, 1996. *supra* and Finn, *et al.*, 1996. *Nucl Acids Res* 24: 3357-3363. For example, a DNA chain can be synthesized on a solid support using standard phosphoramidite coupling chemistry, and modified nucleoside analogs, *e.g.*, 5'-(4-methoxytrityl)amino-5'-deoxy-thymidine phosphoramidite, can be used between the

10   PNA and the 5' end of DNA. *See, e.g.*, Mag, *et al.*, 1989. *Nucl Acid Res* 17: 5973-5988. PNA monomers are then coupled in a stepwise manner to produce a chimeric molecule with a 5' PNA segment and a 3' DNA segment. *See, e.g.*, Finn, *et al.*, 1996. *supra*. Alternatively, chimeric molecules can be synthesized with a 5' DNA segment and a 3' PNA segment. *See, e.g.*, Petersen, *et al.*, 1975. *Bioorg. Med. Chem. Lett.* 5: 1119-11124.

15   In other embodiments, the oligonucleotide may include other appended groups such as peptides (*e.g.*, for targeting host cell receptors *in vivo*), or agents facilitating transport across the cell membrane (*see, e.g.*, Letsinger, *et al.*, 1989. *Proc. Natl. Acad. Sci. U.S.A.* 86: 6553-6556; Lemaitre, *et al.*, 1987. *Proc. Natl. Acad. Sci.* 84: 648-652; PCT Publication No. WO88/09810) or the blood-brain barrier (*see, e.g.*, PCT Publication No. WO 89/10134). In

20   addition, oligonucleotides can be modified with hybridization triggered cleavage agents (*see, e.g.*, Krol, *et al.*, 1988. *BioTechniques* 6:958-976) or intercalating agents (*see, e.g.*, Zon, 1988. *Pharm. Res.* 5: 539-549). To this end, the oligonucleotide may be conjugated to another molecule, *e.g.*, a peptide, a hybridization triggered cross-linking agent, a transport agent, a hybridization-triggered cleavage agent, and the like.

25   **NOVX Polypeptides**

A polypeptide according to the invention includes a polypeptide including the amino acid sequence of NOVX polypeptides whose sequences are provided in any one of SEQ ID NO:2*n*, wherein *n* is an integer between 1 and 124. The invention also includes a mutant or variant protein any of whose residues may be changed from the corresponding

30   residues shown in any one of SEQ ID NO:2*n*, wherein *n* is an integer between 1 and 124, while still encoding a protein that maintains its NOVX activities and physiological functions, or a functional fragment thereof.

In general, a NOVX variant that preserves NOVX-like function includes any variant in which residues at a particular position in the sequence have been substituted by other amino acids, and further include the possibility of inserting an additional residue or residues between two residues of the parent protein as well as the possibility of deleting

5   one or more residues from the parent sequence. Any amino acid substitution, insertion, or deletion is encompassed by the invention. In favorable circumstances, the substitution is a conservative substitution as defined above.

One aspect of the invention pertains to isolated NOVX proteins, and biologically-active portions thereof, or derivatives, fragments, analogs or homologs thereof.

10   Also provided are polypeptide fragments suitable for use as immunogens to raise anti-NOVX antibodies. In one embodiment, native NOVX proteins can be isolated from cells or tissue sources by an appropriate purification scheme using standard protein purification techniques. In another embodiment, NOVX proteins are produced by recombinant DNA techniques. Alternative to recombinant expression, a NOVX protein or

15   polypeptide can be synthesized chemically using standard peptide synthesis techniques.

An "isolated" or "purified" polypeptide or protein or biologically-active portion thereof is substantially free of cellular material or other contaminating proteins from the cell or tissue source from which the NOVX protein is derived, or substantially free from chemical precursors or other chemicals when chemically synthesized. The language

20   "substantially free of cellular material" includes preparations of NOVX proteins in which the protein is separated from cellular components of the cells from which it is isolated or recombinantly-produced. In one embodiment, the language "substantially free of cellular material" includes preparations of NOVX proteins having less than about 30% (by dry weight) of non-NOVX proteins (also referred to herein as a "contaminating protein"), more

25   preferably less than about 20% of non-NOVX proteins, still more preferably less than about 10% of non-NOVX proteins, and most preferably less than about 5% of non-NOVX proteins. When the NOVX protein or biologically-active portion thereof is recombinantly-produced, it is also preferably substantially free of culture medium, i.e., culture medium represents less than about 20%, more preferably less than about 10%, and

30   most preferably less than about 5% of the volume of the NOVX protein preparation.

The language "substantially free of chemical precursors or other chemicals" includes preparations of NOVX proteins in which the protein is separated from chemical precursors or other chemicals that are involved in the synthesis of the protein. In one

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embodiment, the language "substantially free of chemical precursors or other chemicals" includes preparations of NOVX proteins having less than about 30% (by dry weight) of chemical precursors or non-NOVX chemicals, more preferably less than about 20% chemical precursors or non-NOVX chemicals, still more preferably less than about 10%

5 chemical precursors or non-NOVX chemicals, and most preferably less than about 5% chemical precursors or non-NOVX chemicals.

Biologically-active portions of NOVX proteins include peptides comprising amino acid sequences sufficiently homologous to or derived from the amino acid sequences of the NOVX proteins (*e.g.*, the amino acid sequence of SEQ ID NO:2n, wherein n is an integer

10 between 1 and 124) that include fewer amino acids than the full-length NOVX proteins, and exhibit at least one activity of a NOVX protein. Typically, biologically-active portions comprise a domain or motif with at least one activity of the NOVX protein. A biologically-active portion of a NOVX protein can be a polypeptide which is, for example, 10, 25, 50, 100 or more amino acid residues in length.

15 Moreover, other biologically-active portions, in which other regions of the protein are deleted, can be prepared by recombinant techniques and evaluated for one or more of the functional activities of a native NOVX protein.

In an embodiment, the NOVX protein has an amino acid sequence of SEQ ID NO:2n, wherein n is an integer between 1 and 124. In other embodiments, the NOVX

20 protein is substantially homologous to SEQ ID NO:2n, wherein n is an integer between 1 and 124, and retains the functional activity of the protein of SEQ ID NO:2n, wherein n is an integer between 1 and 124, yet differs in amino acid sequence due to natural allelic variation or mutagenesis, as described in detail, below. Accordingly, in another embodiment, the NOVX protein is a protein that comprises an amino acid sequence at least

25 about 45% homologous to the amino acid sequence of SEQ ID NO:2n, wherein n is an integer between 1 and 124, and retains the functional activity of the NOVX proteins of SEQ ID NO:2n, wherein n is an integer between 1 and 124.

Determining Homology Between Two or More Sequences

To determine the percent homology of two amino acid sequences or of two nucleic

30 acids, the sequences are aligned for optimal comparison purposes (*e.g.*, gaps can be introduced in the sequence of a first amino acid or nucleic acid sequence for optimal alignment with a second amino or nucleic acid sequence). The amino acid residues or nucleotides at corresponding amino acid positions or nucleotide positions are then

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compared. When a position in the first sequence is occupied by the same amino acid residue or nucleotide as the corresponding position in the second sequence, then the molecules are homologous at that position (*i.e.*, as used herein amino acid or nucleic acid "homology" is equivalent to amino acid or nucleic acid "identity").

5   The nucleic acid sequence homology may be determined as the degree of identity between two sequences. The homology may be determined using computer programs known in the art, such as GAP software provided in the GCG program package. *See,* Needleman and Wunsch, 1970. *J Mol Biol* 48: 443-453. Using GCG GAP software with the following settings for nucleic acid sequence comparison: GAP creation penalty of 5.0

10   and GAP extension penalty of 0.3, the coding region of the analogous nucleic acid sequences referred to above exhibits a degree of identity preferably of at least 70%, 75%, 80%, 85%, 90%, 95%, 98%, or 99%, with the CDS (encoding) part of the DNA sequence of SEQ ID NO:2n-1, wherein n is an integer between 1 and 124.

The term "sequence identity" refers to the degree to which two polynucleotide or

15   polypeptide sequences are identical on a residue-by-residue basis over a particular region of comparison. The term "percentage of sequence identity" is calculated by comparing two optimally aligned sequences over that region of comparison, determining the number of positions at which the identical nucleic acid base (*e.g.*, A, T, C, G, U, or I, in the case of nucleic acids) occurs in both sequences to yield the number of matched positions, dividing

20   the number of matched positions by the total number of positions in the region of comparison (*i.e.*, the window size), and multiplying the result by 100 to yield the percentage of sequence identity. The term "substantial identity" as used herein denotes a characteristic of a polynucleotide sequence, wherein the polynucleotide comprises a sequence that has at least 80 percent sequence identity, preferably at least 85 percent

25   identity and often 90 to 95 percent sequence identity, more usually at least 99 percent sequence identity as compared to a reference sequence over a comparison region.

Chimeric and Fusion Proteins

The invention also provides NOVX chimeric or fusion proteins. As used herein, a NOVX "chimeric protein" or "fusion protein" comprises a NOVX polypeptide

30   operatively-linked to a non-NOVX polypeptide. An "NOVX polypeptide" refers to a polypeptide having an amino acid sequence corresponding to a NOVX protein of SEQ ID NO:2n, wherein n is an integer between 1 and 124, whereas a "non-NOVX polypeptide" refers to a polypeptide having an amino acid sequence corresponding to a protein that is not

substantially homologous to the NOVX protein, *e.g.*, a protein that is different from the NOVX protein and that is derived from the same or a different organism. Within a NOVX fusion protein the NOVX polypeptide can correspond to all or a portion of a NOVX protein. In one embodiment, a NOVX fusion protein comprises at least one

5 biologically-active portion of a NOVX protein. In another embodiment, a NOVX fusion protein comprises at least two biologically-active portions of a NOVX protein. In yet another embodiment, a NOVX fusion protein comprises at least three biologically-active portions of a NOVX protein. Within the fusion protein, the term "operatively-linked" is intended to indicate that the NOVX polypeptide and the non-NOVX polypeptide are fused

10 in-frame with one another. The non-NOVX polypeptide can be fused to the N-terminus or C-terminus of the NOVX polypeptide.

In one embodiment, the fusion protein is a GST-NOVX fusion protein in which the NOVX sequences are fused to the C-terminus of the GST (glutathione S-transferase) sequences. Such fusion proteins can facilitate the purification of recombinant NOVX

15 polypeptides.

In another embodiment, the fusion protein is a NOVX protein containing a heterologous signal sequence at its N-terminus. In certain host cells (*e.g.*, mammalian host cells), expression and/or secretion of NOVX can be increased through use of a heterologous signal sequence.

20 In yet another embodiment, the fusion protein is a NOVX-immunoglobulin fusion protein in which the NOVX sequences are fused to sequences derived from a member of the immunoglobulin protein family. The NOVX-immunoglobulin fusion proteins of the invention can be incorporated into pharmaceutical compositions and administered to a subject to inhibit an interaction between a NOVX ligand and a NOVX protein on the

25 surface of a cell, to thereby suppress NOVX-mediated signal transduction *in vivo*. The NOVX-immunoglobulin fusion proteins can be used to affect the bioavailability of a NOVX cognate ligand. Inhibition of the NOVX ligand/NOVX interaction may be useful therapeutically for both the treatment of proliferative and differentiative disorders, as well as modulating (*e.g.* promoting or inhibiting) cell survival. Moreover, the

30 NOVX-immunoglobulin fusion proteins of the invention can be used as immunogens to produce anti-NOVX antibodies in a subject, to purify NOVX ligands, and in screening assays to identify molecules that inhibit the interaction of NOVX with a NOVX ligand.

A NOVX chimeric or fusion protein of the invention can be produced by standard recombinant DNA techniques. For example, DNA fragments coding for the different polypeptide sequences are ligated together in-frame in accordance with conventional techniques, *e.g.*, by employing blunt-ended or stagger-ended termini for ligation, restriction

5   enzyme digestion to provide for appropriate termini, filling-in of cohesive ends as appropriate, alkaline phosphatase treatment to avoid undesirable joining, and enzymatic ligation. In another embodiment, the fusion gene can be synthesized by conventional techniques including automated DNA synthesizers. Alternatively, PCR amplification of gene fragments can be carried out using anchor primers that give rise to complementary

10   overhangs between two consecutive gene fragments that can subsequently be annealed and reamplified to generate a chimeric gene sequence (*see, e.g.*, Ausubel, *et al.* (eds.) CURRENT PROTOCOLS IN MOLECULAR BIOLOGY, John Wiley & Sons, 1992). Moreover, many expression vectors are commercially available that already encode a fusion moiety (*e.g.*, a GST polypeptide). A NOVX-encoding nucleic acid can be cloned into such an expression

15   vector such that the fusion moiety is linked in-frame to the NOVX protein.

NOVX Agonists and Antagonists

The invention also pertains to variants of the NOVX proteins that function as either NOVX agonists (*i.e.*, mimetics) or as NOVX antagonists. Variants of the NOVX protein can be generated by mutagenesis (*e.g.*, discrete point mutation or truncation of the NOVX

20   protein). An agonist of the NOVX protein can retain substantially the same, or a subset of, the biological activities of the naturally occurring form of the NOVX protein. An antagonist of the NOVX protein can inhibit one or more of the activities of the naturally occurring form of the NOVX protein by, for example, competitively binding to a downstream or upstream member of a cellular signaling cascade which includes the NOVX

25   protein. Thus, specific biological effects can be elicited by treatment with a variant of limited function. In one embodiment, treatment of a subject with a variant having a subset of the biological activities of the naturally occurring form of the protein has fewer side effects in a subject relative to treatment with the naturally occurring form of the NOVX proteins.

30   Variants of the NOVX proteins that function as either NOVX agonists (*i.e.*, mimetics) or as NOVX antagonists can be identified by screening combinatorial libraries of mutants (*e.g.*, truncation mutants) of the NOVX proteins for NOVX protein agonist or antagonist activity. In one embodiment, a variegated library of NOVX variants is

generated by combinatorial mutagenesis at the nucleic acid level and is encoded by a variegated gene library. A variegated library of NOVX variants can be produced by, for example, enzymatically ligating a mixture of synthetic oligonucleotides into gene sequences such that a degenerate set of potential NOVX sequences is expressible as individual polypeptides, or alternatively, as a set of larger fusion proteins (*e.g.*, for phage display) containing the set of NOVX sequences therein. There are a variety of methods which can be used to produce libraries of potential NOVX variants from a degenerate oligonucleotide sequence. Chemical synthesis of a degenerate gene sequence can be performed in an automatic DNA synthesizer, and the synthetic gene then ligated into an appropriate expression vector. Use of a degenerate set of genes allows for the provision, in one mixture, of all of the sequences encoding the desired set of potential NOVX sequences. Methods for synthesizing degenerate oligonucleotides are well-known within the art. *See, e.g.*, Narang, 1983. *Tetrahedron* 39: 3; Itakura, *et al.*, 1984. *Annu. Rev. Biochem.* 53: 323; Itakura, *et al.*, 1984. *Science* 198: 1056; Ike, *et al.*, 1983. *Nucl. Acids Res.* 11: 477.

Polypeptide Libraries

In addition, libraries of fragments of the NOVX protein coding sequences can be used to generate a variegated population of NOVX fragments for screening and subsequent selection of variants of a NOVX protein. In one embodiment, a library of coding sequence fragments can be generated by treating a double stranded PCR fragment of a NOVX coding sequence with a nuclease under conditions wherein nicking occurs only about once per molecule, denaturing the double stranded DNA, renaturing the DNA to form double-stranded DNA that can include sense/antisense pairs from different nicked products, removing single stranded portions from reformed duplexes by treatment with S_1 nuclease, and ligating the resulting fragment library into an expression vector. By this method, expression libraries can be derived which encodes N-terminal and internal fragments of various sizes of the NOVX proteins.

Various techniques are known in the art for screening gene products of combinatorial libraries made by point mutations or truncation, and for screening cDNA libraries for gene products having a selected property. Such techniques are adaptable for rapid screening of the gene libraries generated by the combinatorial mutagenesis of NOVX proteins. The most widely used techniques, which are amenable to high throughput analysis, for screening large gene libraries typically include cloning the gene library into replicable expression vectors, transforming appropriate cells with the resulting library of

vectors, and expressing the combinatorial genes under conditions in which detection of a desired activity facilitates isolation of the vector encoding the gene whose product was detected. Recursive ensemble mutagenesis (REM), a new technique that enhances the frequency of functional mutants in the libraries, can be used in combination with the

5 screening assays to identify NOVX variants. *See, e.g.,* Arkin and Yourvan, 1992. *Proc. Natl. Acad. Sci. USA* 89: 7811-7815; Delgrave, *et al.,* 1993. *Protein Engineering* 6:327-331.

Anti-NOVX Antibodies

Included in the invention are antibodies to NOVX proteins, or fragments of NOVX

10 proteins. The term "antibody" as used herein refers to immunoglobulin molecules and immunologically active portions of immunoglobulin (Ig) molecules, *i.e.,* molecules that contain an antigen binding site that specifically binds (immunoreacts with) an antigen. Such antibodies include, but are not limited to, polyclonal, monoclonal, chimeric, single chain, F_{ab}, $F_{ab'}$ and $F_{(ab')2}$ fragments, and an F_{ab} expression library. In general, antibody

15 molecules obtained from humans relates to any of the classes IgG, IgM, IgA, IgE and IgD, which differ from one another by the nature of the heavy chain present in the molecule. Certain classes have subclasses as well, such as IgG_1, IgG_2, and others. Furthermore, in humans, the light chain may be a kappa chain or a lambda chain. Reference herein to antibodies includes a reference to all such classes, subclasses and types of human antibody

20 species.

An isolated protein of the invention intended to serve as an antigen, or a portion or fragment thereof, can be used as an immunogen to generate antibodies that immunospecifically bind the antigen, using standard techniques for polyclonal and monoclonal antibody preparation. The full-length protein can be used or, alternatively, the

25 invention provides antigenic peptide fragments of the antigen for use as immunogens. An antigenic peptide fragment comprises at least 6 amino acid residues of the amino acid sequence of the full length protein, such as an amino acid sequence of SEQ ID NO:$2n$, wherein n is an integer between 1 and 124, and encompasses an epitope thereof such that an antibody raised against the peptide forms a specific immune complex with the full

30 length protein or with any fragment that contains the epitope. Preferably, the antigenic peptide comprises at least 10 amino acid residues, or at least 15 amino acid residues, or at least 20 amino acid residues, or at least 30 amino acid residues. Preferred epitopes

encompassed by the antigenic peptide are regions of the protein that are located on its surface; commonly these are hydrophilic regions.

In certain embodiments of the invention, at least one epitope encompassed by the antigenic peptide is a region of NOVX that is located on the surface of the protein, *e.g.*, a hydrophilic region. A hydrophobicity analysis of the human NOVX protein sequence will indicate which regions of a NOVX polypeptide are particularly hydrophilic and, therefore, are likely to encode surface residues useful for targeting antibody production. As a means for targeting antibody production, hydropathy plots showing regions of hydrophilicity and hydrophobicity may be generated by any method well known in the art, including, for example, the Kyte Doolittle or the Hopp Woods methods, either with or without Fourier transformation. See, *e.g.*, Hopp and Woods, 1981, *Proc. Nat. Acad. Sci. USA* 78: 3824-3828; Kyte and Doolittle 1982, *J. Mol. Biol.* 157: 105-142, each incorporated herein by reference in their entirety. Antibodies that are specific for one or more domains within an antigenic protein, or derivatives, fragments, analogs or homologs thereof, are also provided herein.

The term "epitope" includes any protein determinant capable of specific binding to an immunoglobulin or T-cell receptor. Epitopic determinants usually consist of chemically active surface groupings of molecules such as amino acids or sugar side chains and usually have specific three dimensional structural characteristics, as well as specific charge characteristics. A NOVX polypeptide or a fragment thereof comprises at least one antigenic epitope. An anti-NOVX antibody of the present invention is said to specifically bind to antigen NOVX when the equilibrium binding constant (K_D) is ≤1 μM, preferably ≤ 100 nM, more preferably ≤ 10 nM, and most preferably ≤ 100 pM to about 1 pM, as measured by assays such as radioligand binding assays or similar assays known to those skilled in the art.

A protein of the invention, or a derivative, fragment, analog, homolog or ortholog thereof, may be utilized as an immunogen in the generation of antibodies that immunospecifically bind these protein components.

Various procedures known within the art may be used for the production of polyclonal or monoclonal antibodies directed against a protein of the invention, or against derivatives, fragments, analogs homologs or orthologs thereof (see, for example, Antibodies: A Laboratory Manual, Harlow E, and Lane D, 1988, Cold Spring Harbor

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Laboratory Press, Cold Spring Harbor, NY, incorporated herein by reference). Some of these antibodies are discussed below.

Polyclonal Antibodies

For the production of polyclonal antibodies, various suitable host animals (*e.g.*,
5   rabbit, goat, mouse or other mammal) may be immunized by one or more injections with the native protein, a synthetic variant thereof, or a derivative of the foregoing. An appropriate immunogenic preparation can contain, for example, the naturally occurring immunogenic protein, a chemically synthesized polypeptide representing the immunogenic protein, or a recombinantly expressed immunogenic protein. Furthermore, the protein may
10   be conjugated to a second protein known to be immunogenic in the mammal being immunized. Examples of such immunogenic proteins include but are not limited to keyhole limpet hemocyanin, serum albumin, bovine thyroglobulin, and soybean trypsin inhibitor. The preparation can further include an adjuvant. Various adjuvants used to increase the immunological response include, but are not limited to, Freund's (complete and
15   incomplete), mineral gels (*e.g.*, aluminum hydroxide), surface active substances (*e.g.*, lysolecithin, pluronic polyols, polyanions, peptides, oil emulsions, dinitrophenol, *etc.*), adjuvants usable in humans such as Bacille Calmette-Guerin and Corynebacterium parvum, or similar immunostimulatory agents. Additional examples of adjuvants which can be employed include MPL-TDM adjuvant (monophosphoryl Lipid A, synthetic trehalose
20   dicorynomycolate).

The polyclonal antibody molecules directed against the immunogenic protein can be isolated from the mammal (*e.g.*, from the blood) and further purified by well known techniques, such as affinity chromatography using protein A or protein G, which provide primarily the IgG fraction of immune serum. Subsequently, or alternatively, the specific
25   antigen which is the target of the immunoglobulin sought, or an epitope thereof, may be immobilized on a column to purify the immune specific antibody by immunoaffinity chromatography. Purification of immunoglobulins is discussed, for example, by D. Wilkinson (The Scientist, published by The Scientist, Inc., Philadelphia PA, Vol. 14, No. 8 (April 17, 2000), pp. 25-28).

30   ## Monoclonal Antibodies

The term "monoclonal antibody" (MAb) or "monoclonal antibody composition", as used herein, refers to a population of antibody molecules that contain only one molecular

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species of antibody molecule consisting of a unique light chain gene product and a unique heavy chain gene product. In particular, the complementarity determining regions (CDRs) of the monoclonal antibody are identical in all the molecules of the population. MAbs thus contain an antigen binding site capable of immunoreacting with a particular epitope of the

5   antigen characterized by a unique binding affinity for it.

       Monoclonal antibodies can be prepared using hybridoma methods, such as those described by Kohler and Milstein, Nature, 256:495 (1975). In a hybridoma method, a mouse, hamster, or other appropriate host animal, is typically immunized with an immunizing agent to elicit lymphocytes that produce or are capable of producing antibodies

10   that will specifically bind to the immunizing agent. Alternatively, the lymphocytes can be immunized in vitro.

       The immunizing agent will typically include the protein antigen, a fragment thereof or a fusion protein thereof. Generally, either peripheral blood lymphocytes are used if cells of human origin are desired, or spleen cells or lymph node cells are used if non-human

15   mammalian sources are desired. The lymphocytes are then fused with an immortalized cell line using a suitable fusing agent, such as polyethylene glycol, to form a hybridoma cell (Goding, Monoclonal Antibodies: Principles and Practice, Academic Press, (1986) pp. 59-103). Immortalized cell lines are usually transformed mammalian cells, particularly myeloma cells of rodent, bovine and human origin. Usually, rat or mouse myeloma cell

20   lines are employed. The hybridoma cells can be cultured in a suitable culture medium that preferably contains one or more substances that inhibit the growth or survival of the unfused, immortalized cells. For example, if the parental cells lack the enzyme hypoxanthine guanine phosphoribosyl transferase (HGPRT or HPRT), the culture medium for the hybridomas typically will include hypoxanthine, aminopterin, and thymidine ("HAT

25   medium"), which substances prevent the growth of HGPRT-deficient cells.

       Preferred immortalized cell lines are those that fuse efficiently, support stable high level expression of antibody by the selected antibody-producing cells, and are sensitive to a medium such as HAT medium. More preferred immortalized cell lines are murine myeloma lines, which can be obtained, for instance, from the Salk Institute Cell

30   Distribution Center, San Diego, California and the American Type Culture Collection, Manassas, Virginia. Human myeloma and mouse-human heteromyeloma cell lines also have been described for the production of human monoclonal antibodies (Kozbor, J.

Immunol., 133:3001 (1984); Brodeur et al., Monoclonal Antibody Production Techniques and Applications, Marcel Dekker, Inc., New York, (1987) pp. 51-63).

The culture medium in which the hybridoma cells are cultured can then be assayed for the presence of monoclonal antibodies directed against the antigen. Preferably, the

5   binding specificity of monoclonal antibodies produced by the hybridoma cells is determined by immunoprecipitation or by an in vitro binding assay, such as radioimmunoassay (RIA) or enzyme-linked immunoabsorbent assay (ELISA). Such techniques and assays are known in the art. The binding affinity of the monoclonal antibody can, for example, be determined by the Scatchard analysis of Munson and Pollard,

10   Anal. Biochem., 107:220 (1980). It is an objective, especially important in therapeutic applications of monoclonal antibodies, to identify antibodies having a high degree of specificity and a high binding affinity for the target antigen.

After the desired hybridoma cells are identified, the clones can be subcloned by limiting dilution procedures and grown by standard methods (Goding,1986). Suitable

15   culture media for this purpose include, for example, Dulbecco's Modified Eagle's Medium and RPMI-1640 medium. Alternatively, the hybridoma cells can be grown in vivo as ascites in a mammal.

The monoclonal antibodies secreted by the subclones can be isolated or purified from the culture medium or ascites fluid by conventional immunoglobulin purification

20   procedures such as, for example, protein A-Sepharose, hydroxylapatite chromatography, gel electrophoresis, dialysis, or affinity chromatography.

The monoclonal antibodies can also be made by recombinant DNA methods, such as those described in U.S. Patent No. 4,816,567. DNA encoding the monoclonal antibodies of the invention can be readily isolated and sequenced using conventional procedures (e.g.,

25   by using oligonucleotide probes that are capable of binding specifically to genes encoding the heavy and light chains of murine antibodies). The hybridoma cells of the invention serve as a preferred source of such DNA. Once isolated, the DNA can be placed into expression vectors, which are then transfected into host cells such as simian COS cells, Chinese hamster ovary (CHO) cells, or myeloma cells that do not otherwise produce

30   immunoglobulin protein, to obtain the synthesis of monoclonal antibodies in the recombinant host cells. The DNA also can be modified, for example, by substituting the coding sequence for human heavy and light chain constant domains in place of the homologous murine sequences (U.S. Patent No. 4,816,567; Morrison, Nature 368, 812-13

(1994)) or by covalently joining to the immunoglobulin coding sequence all or part of the coding sequence for a non-immunoglobulin polypeptide. Such a non-immunoglobulin polypeptide can be substituted for the constant domains of an antibody of the invention, or can be substituted for the variable domains of one antigen-combining site of an antibody of the invention to create a chimeric bivalent antibody.

Humanized Antibodies

The antibodies directed against the protein antigens of the invention can further comprise humanized antibodies or human antibodies. These antibodies are suitable for administration to humans without engendering an immune response by the human against the administered immunoglobulin. Humanized forms of antibodies are chimeric immunoglobulins, immunoglobulin chains or fragments thereof (such as Fv, Fab, Fab', F(ab')$_2$ or other antigen-binding subsequences of antibodies) that are principally comprised of the sequence of a human immunoglobulin, and contain minimal sequence derived from a non-human immunoglobulin. Humanization can be performed following the method of Winter and co-workers (Jones et al., Nature, 321:522-525 (1986); Riechmann et al., Nature, 332:323-327 (1988); Verhoeyen et al., Science, 239:1534-1536 (1988)), by substituting rodent CDRs or CDR sequences for the corresponding sequences of a human antibody. (See also U.S. Patent No. 5,225,539.) In some instances, Fv framework residues of the human immunoglobulin are replaced by corresponding non-human residues. Humanized antibodies can also comprise residues which are found neither in the recipient antibody nor in the imported CDR or framework sequences. In general, the humanized antibody will comprise substantially all of at least one, and typically two, variable domains, in which all or substantially all of the CDR regions correspond to those of a non-human immunoglobulin and all or substantially all of the framework regions are those of a human immunoglobulin consensus sequence. The humanized antibody optimally also will comprise at least a portion of an immunoglobulin constant region (Fc), typically that of a human immunoglobulin (Jones et al., 1986; Riechmann et al., 1988; and Presta, Curr. Op. Struct. Biol., 2:593-596 (1992)).

Human Antibodies

Fully human antibodies essentially relate to antibody molecules in which the entire sequence of both the light chain and the heavy chain, including the CDRs, arise from human genes. Such antibodies are termed "human antibodies", or "fully human antibodies"

herein. Human monoclonal antibodies can be prepared by the trioma technique; the human B-cell hybridoma technique (see Kozbor, et al., 1983 Immunol Today 4: 72) and the EBV hybridoma technique to produce human monoclonal antibodies (see Cole, et al., 1985 In: MONOCLONAL ANTIBODIES AND CANCER THERAPY, Alan R. Liss, Inc., pp. 77-96). Human

5   monoclonal antibodies may be utilized in the practice of the present invention and may be produced by using human hybridomas (see Cote, et al., 1983. Proc Natl Acad Sci USA 80: 2026-2030) or by transforming human B-cells with Epstein Barr Virus in vitro (see Cole, et al., 1985 In: MONOCLONAL ANTIBODIES AND CANCER THERAPY, Alan R. Liss, Inc., pp. 77-96).

10   In addition, human antibodies can also be produced using additional techniques, including phage display libraries (Hoogenboom and Winter, J. Mol. Biol., 227:381 (1991); Marks et al., J. Mol. Biol., 222:581 (1991)). Similarly, human antibodies can be made by introducing human immunoglobulin loci into transgenic animals, e.g., mice in which the endogenous immunoglobulin genes have been partially or completely inactivated. Upon

15   challenge, human antibody production is observed, which closely resembles that seen in humans in all respects, including gene rearrangement, assembly, and antibody repertoire. This approach is described, for example, in U.S. Patent Nos. 5,545,807; 5,545,806; 5,569,825; 5,625,126; 5,633,425; 5,661,016, and in Marks et al. (Bio/Technology 10, 779-783 (1992)); Lonberg et al. (Nature 368 856-859 (1994)); Morrison (Nature 368,

20   812-13 (1994)); Fishwild et al,(Nature Biotechnology 14, 845-51 (1996)); Neuberger (Nature Biotechnology 14, 826 (1996)); and Lonberg and Huszar (Intern. Rev. Immunol. 13 65-93 (1995)).

Human antibodies may additionally be produced using transgenic nonhuman animals which are modified so as to produce fully human antibodies rather than the

25   animal's endogenous antibodies in response to challenge by an antigen. (See PCT publication WO94/02602). The endogenous genes encoding the heavy and light immunoglobulin chains in the nonhuman host have been incapacitated, and active loci encoding human heavy and light chain immunoglobulins are inserted into the host's genome. The human genes are incorporated, for example, using yeast artificial

30   chromosomes containing the requisite human DNA segments. An animal which provides all the desired modifications is then obtained as progeny by crossbreeding intermediate transgenic animals containing fewer than the full complement of the modifications. The preferred embodiment of such a nonhuman animal is a mouse, and is termed the

Xenomouse™ as disclosed in PCT publications WO 96/33735 and WO 96/34096. This animal produces B cells which secrete fully human immunoglobulins. The antibodies can be obtained directly from the animal after immunization with an immunogen of interest, as, for example, a preparation of a polyclonal antibody, or alternatively from immortalized B

5   cells derived from the animal, such as hybridomas producing monoclonal antibodies. Additionally, the genes encoding the immunoglobulins with human variable regions can be recovered and expressed to obtain the antibodies directly, or can be further modified to obtain analogs of antibodies such as, for example, single chain Fv molecules.

An example of a method of producing a nonhuman host, exemplified as a mouse,

10   lacking expression of an endogenous immunoglobulin heavy chain is disclosed in U.S. Patent No. 5,939,598. It can be obtained by a method including deleting the J segment genes from at least one endogenous heavy chain locus in an embryonic stem cell to prevent rearrangement of the locus and to prevent formation of a transcript of a rearranged immunoglobulin heavy chain locus, the deletion being effected by a targeting vector

15   containing a gene encoding a selectable marker; and producing from the embryonic stem cell a transgenic mouse whose somatic and germ cells contain the gene encoding the selectable marker.

A method for producing an antibody of interest, such as a human antibody, is disclosed in U.S. Patent No. 5,916,771. It includes introducing an expression vector that

20   contains a nucleotide sequence encoding a heavy chain into one mammalian host cell in culture, introducing an expression vector containing a nucleotide sequence encoding a light chain into another mammalian host cell, and fusing the two cells to form a hybrid cell. The hybrid cell expresses an antibody containing the heavy chain and the light chain.

In a further improvement on this procedure, a method for identifying a clinically

25   relevant epitope on an immunogen, and a correlative method for selecting an antibody that binds immunospecifically to the relevant epitope with high affinity, are disclosed in PCT publication WO 99/53049.

F_{ab} Fragments and Single Chain Antibodies

According to the invention, techniques can be adapted for the production of

30   single-chain antibodies specific to an antigenic protein of the invention (see e.g., U.S. Patent No. 4,946,778). In addition, methods can be adapted for the construction of F_{ab} expression libraries (see e.g., Huse, et al., 1989 Science 246: 1275-1281) to allow rapid and effective identification of monoclonal F_{ab} fragments with the desired specificity for a

protein or derivatives, fragments, analogs or homologs thereof. Antibody fragments that

contain the idiotypes to a protein antigen may be produced by techniques known in the art

including, but not limited to: (i) an $F_{(ab')2}$ fragment produced by pepsin digestion of an

antibody molecule; (ii) an F_{ab} fragment generated by reducing the disulfide bridges of an

5      $F_{(ab')2}$ fragment; (iii) an F_{ab} fragment generated by the treatment of the antibody molecule

with papain and a reducing agent and (iv) F_v fragments.

Bispecific Antibodies

Bispecific antibodies are monoclonal, preferably human or humanized, antibodies

that have binding specificities for at least two different antigens. In the present case, one of

10   the binding specificities is for an antigenic protein of the invention. The second binding

target is any other antigen, and advantageously is a cell-surface protein or receptor or

receptor subunit.

Methods for making bispecific antibodies are known in the art. Traditionally, the

recombinant production of bispecific antibodies is based on the co-expression of two

15   immunoglobulin heavy-chain/light-chain pairs, where the two heavy chains have different

specificities (Milstein and Cuello, Nature, 305:537-539 (1983)). Because of the random

assortment of immunoglobulin heavy and light chains, these hybridomas (quadromas)

produce a potential mixture of ten different antibody molecules, of which only one has the

correct bispecific structure. The purification of the correct molecule is usually

20   accomplished by affinity chromatography steps. Similar procedures are disclosed in WO

93/08829, published 13 May 1993, and in Traunecker et al., EMBO J., 10:3655-3659

(1991).

Antibody variable domains with the desired binding specificities (antibody-antigen

combining sites) can be fused to immunoglobulin constant domain sequences. The fusion

25   preferably is with an immunoglobulin heavy-chain constant domain, comprising at least

part of the hinge, CH2, and CH3 regions. It is preferred to have the first heavy-chain

constant region (CH1) containing the site necessary for light-chain binding present in at

least one of the fusions. DNAs encoding the immunoglobulin heavy-chain fusions and, if

desired, the immunoglobulin light chain, are inserted into separate expression vectors, and

30   are co-transfected into a suitable host organism. For further details of generating bispecific

antibodies see, for example, Suresh et al., Methods in Enzymology, 121:210 (1986).

According to another approach described in WO 96/27011, the interface between a

pair of antibody molecules can be engineered to maximize the percentage of heterodimers

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which are recovered from recombinant cell culture. The preferred interface comprises at least a part of the CH3 region of an antibody constant domain. In this method, one or more small amino acid side chains from the interface of the first antibody molecule are replaced with larger side chains (*e.g.* tyrosine or tryptophan). Compensatory "cavities" of identical

5   or similar size to the large side chain(s) are created on the interface of the second antibody molecule by replacing large amino acid side chains with smaller ones (*e.g.* alanine or threonine). This provides a mechanism for increasing the yield of the heterodimer over other unwanted end-products such as homodimers.

Bispecific antibodies can be prepared as full length antibodies or antibody

10   fragments (*e.g.* F(ab')$_2$ bispecific antibodies). Techniques for generating bispecific antibodies from antibody fragments have been described in the literature. For example, bispecific antibodies can be prepared using chemical linkage. Brennan et al., Science 229:81 (1985) describe a procedure wherein intact antibodies are proteolytically cleaved to generate F(ab')$_2$ fragments. These fragments are reduced in the presence of the dithiol

15   complexing agent sodium arsenite to stabilize vicinal dithiols and prevent intermolecular disulfide formation. The Fab' fragments generated are then converted to thionitrobenzoate (TNB) derivatives. One of the Fab'-TNB derivatives is then reconverted to the Fab'-thiol by reduction with mercaptoethylamine and is mixed with an equimolar amount of the other Fab'-TNB derivative to form the bispecific antibody. The bispecific antibodies produced

20   can be used as agents for the selective immobilization of enzymes.

Additionally, Fab' fragments can be directly recovered from E. coli and chemically coupled to form bispecific antibodies. Shalaby et al., J. Exp. Med. 175:217-225 (1992) describe the production of a fully humanized bispecific antibody F(ab')$_2$ molecule. Each Fab' fragment was separately secreted from E. coli and subjected to directed chemical

25   coupling in vitro to form the bispecific antibody. The bispecific antibody thus formed was able to bind to cells overexpressing the ErbB2 receptor and normal human T cells, as well as trigger the lytic activity of human cytotoxic lymphocytes against human breast tumor targets.

Various techniques for making and isolating bispecific antibody fragments directly

30   from recombinant cell culture have also been described. For example, bispecific antibodies have been produced using leucine zippers. Kostelny et al., J. Immunol. 148(5):1547-1553 (1992). The leucine zipper peptides from the Fos and Jun proteins were linked to the Fab' portions of two different antibodies by gene fusion. The antibody homodimers were

reduced at the hinge region to form monomers and then re-oxidized to form the antibody heterodimers. This method can also be utilized for the production of antibody homodimers. The "diabody" technology described by Hollinger et al., Proc. Natl. Acad. Sci. USA 90:6444-6448 (1993) has provided an alternative mechanism for making bispecific antibody fragments. The fragments comprise a heavy-chain variable domain (V_H) connected to a light-chain variable domain (V_L) by a linker which is too short to allow pairing between the two domains on the same chain. Accordingly, the V_H and V_L domains of one fragment are forced to pair with the complementary V_L and V_H domains of another fragment, thereby forming two antigen-binding sites. Another strategy for making bispecific antibody fragments by the use of single-chain Fv (sFv) dimers has also been reported. See, Gruber et al., J. Immunol. 152:5368 (1994).

Antibodies with more than two valencies are contemplated. For example, trispecific antibodies can be prepared. Tutt et al., J. Immunol. 147:60 (1991).

Exemplary bispecific antibodies can bind to two different epitopes, at least one of which originates in the protein antigen of the invention. Alternatively, an anti-antigenic arm of an immunoglobulin molecule can be combined with an arm which binds to a triggering molecule on a leukocyte such as a T-cell receptor molecule (*e.g.* CD2, CD3, CD28, or B7), or Fc receptors for IgG (FcγR), such as FcγRI (CD64), FcγRII (CD32) and FcγRIII (CD16) so as to focus cellular defense mechanisms to the cell expressing the particular antigen. Bispecific antibodies can also be used to direct cytotoxic agents to cells which express a particular antigen. These antibodies possess an antigen-binding arm and an arm which binds a cytotoxic agent or a radionuclide chelator, such as EOTUBE, DPTA, DOTA, or TETA. Another bispecific antibody of interest binds the protein antigen described herein and further binds tissue factor (TF).

Heteroconjugate Antibodies

Heteroconjugate antibodies are also within the scope of the present invention. Heteroconjugate antibodies are composed of two covalently joined antibodies. Such antibodies have, for example, been proposed to target immune system cells to unwanted cells (U.S. Patent No. 4,676,980), and for treatment of HIV infection (WO 91/00360; WO 92/200373; EP 03089). It is contemplated that the antibodies can be prepared in vitro using known methods in synthetic protein chemistry, including those involving crosslinking agents. For example, immunotoxins can be constructed using a disulfide exchange reaction or by forming a thioether bond. Examples of suitable reagents for this purpose include

iminothiolate and methyl-4-mercaptobutyrimidate and those disclosed, for example, in U.S. Patent No. 4,676,980.

Effector Function Engineering

It can be desirable to modify the antibody of the invention with respect to effector

5   function, so as to enhance, *e.g.*, the effectiveness of the antibody in treating cancer. For example, cysteine residue(s) can be introduced into the Fc region, thereby allowing interchain disulfide bond formation in this region. The homodimeric antibody thus generated can have improved internalization capability and/or increased complement-mediated cell killing and antibody-dependent cellular cytotoxicity (ADCC).

10   See Caron et al., J. Exp Med., 176: 1191-1195 (1992) and Shopes, J. Immunol., 148: 2918-2922 (1992). Homodimeric antibodies with enhanced anti-tumor activity can also be prepared using heterobifunctional cross-linkers as described in Wolff et al. Cancer Research, 53: 2560-2565 (1993). Alternatively, an antibody can be engineered that has dual Fc regions and can thereby have enhanced complement lysis and ADCC capabilities.

15   See Stevenson et al., Anti-Cancer Drug Design, 3: 219-230 (1989).

Immunoconjugates

The invention also pertains to immunoconjugates comprising an antibody conjugated to a cytotoxic agent such as a chemotherapeutic agent, toxin (*e.g.*, an enzymatically active toxin of bacterial, fungal, plant, or animal origin, or fragments

20   thereof), or a radioactive isotope (*i.e.*, a radioconjugate).

Chemotherapeutic agents useful in the generation of such immunoconjugates have been described above. Enzymatically active toxins and fragments thereof that can be used include diphtheria A chain, nonbinding active fragments of diphtheria toxin, exotoxin A chain (from Pseudomonas aeruginosa), ricin A chain, abrin A chain, modeccin A chain,

25   alpha-sarcin, Aleurites fordii proteins, dianthin proteins, Phytolaca americana proteins (PAPI, PAPII, and PAP-S), momordica charantia inhibitor, curcin, crotin, sapaonaria officinalis inhibitor, gelonin, mitogellin, restrictocin, phenomycin, enomycin, and the tricothecenes. A variety of radionuclides are available for the production of radioconjugated antibodies. Examples include ^{212}Bi, ^{131}I, ^{131}In, ^{90}Y, and ^{186}Re.

30   Conjugates of the antibody and cytotoxic agent are made using a variety of bifunctional protein-coupling agents such as N-succinimidyl-3-(2-pyridyldithiol) propionate (SPDP), iminothiolane (IT), bifunctional derivatives of imidoesters (such as

dimethyl adipimidate HCL), active esters (such as disuccinimidyl suberate), aldehydes
(such as glutareldehyde), bis-azido compounds (such as bis (p-azidobenzoyl)
hexanediamine), bis-diazonium derivatives (such as
bis-(p-diazoniumbenzoyl)-ethylenediamine), diisocyanates (such as tolyene

5   2,6-diisocyanate), and bis-active fluorine compounds (such as
1,5-difluoro-2,4-dinitrobenzene). For example, a ricin immunotoxin can be prepared as
described in Vitetta et al., Science, 238: 1098 (1987). Carbon-14-labeled
1-isothiocyanatobenzyl-3-methyldiethylene triaminepentaacetic acid (MX-DTPA) is an
exemplary chelating agent for conjugation of radionucleotide to the antibody. See

10   WO94/11026.

In another embodiment, the antibody can be conjugated to a "receptor" (such
streptavidin) for utilization in tumor pretargeting wherein the antibody-receptor conjugate
is administered to the patient, followed by removal of unbound conjugate from the
circulation using a clearing agent and then administration of a "ligand" (e.g., avidin) that is

15   in turn conjugated to a cytotoxic agent.

Immunoliposomes

The antibodies disclosed herein can also be formulated as immunoliposomes.
Liposomes containing the antibody are prepared by methods known in the art, such as
described in Epstein et al., Proc. Natl. Acad. Sci. USA, 82: 3688 (1985); Hwang et al.,

20   Proc. Natl Acad. Sci. USA, 77: 4030 (1980); and U.S. Pat. Nos. 4,485,045 and 4,544,545.
Liposomes with enhanced circulation time are disclosed in U.S. Patent No. 5,013,556.

Particularly useful liposomes can be generated by the reverse-phase evaporation
method with a lipid composition comprising phosphatidylcholine, cholesterol, and
PEG-derivatized phosphatidylethanolamine (PEG-PE). Liposomes are extruded through

25   filters of defined pore size to yield liposomes with the desired diameter. Fab' fragments of
the antibody of the present invention can be conjugated to the liposomes as described in
Martin et al., J. Biol. Chem., 257: 286-288 (1982) via a disulfide-interchange reaction. A
chemotherapeutic agent (such as Doxorubicin) is optionally contained within the liposome.
See Gabizon et al., J. National Cancer Inst., 81(19): 1484 (1989).

30   ### Diagnostic Applications of Antibodies Directed Against the Proteins of the Invention

In one embodiment, methods for the screening of antibodies that possess the desired
specificity include, but are not limited to, enzyme linked immunosorbent assay (ELISA)

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and other immunologically mediated techniques known within the art. In a specific embodiment, selection of antibodies that are specific to a particular domain of an NOVX protein is facilitated by generation of hybridomas that bind to the fragment of an NOVX protein possessing such a domain. Thus, antibodies that are specific for a desired domain

5 within an NOVX protein, or derivatives, fragments, analogs or homologs thereof, are also provided herein.

Antibodies directed against a NOVX protein of the invention may be used in methods known within the art relating to the localization and/or quantitation of a NOVX protein (*e.g.*, for use in measuring levels of the NOVX protein within appropriate

10 physiological samples, for use in diagnostic methods, for use in imaging the protein, and the like). In a given embodiment, antibodies specific to a NOVX protein, or derivative, fragment, analog or homolog thereof, that contain the antibody derived antigen binding domain, are utilized as pharmacologically active compounds (referred to hereinafter as "Therapeutics").

15 An antibody specific for a NOVX protein of the invention (e.g., a monoclonal antibody or a polyclonal antibody) can be used to isolate a NOVX polypeptide by standard techniques, such as immunoaffinity, chromatography or immunoprecipitation. An antibody to a NOVX polypeptide can facilitate the purification of a natural NOVX antigen from cells, or of a recombinantly produced NOVX antigen expressed in host cells. Moreover,

20 such an anti-NOVX antibody can be used to detect the antigenic NOVX protein (*e.g.*, in a cellular lysate or cell supernatant) in order to evaluate the abundance and pattern of expression of the antigenic NOVX protein. Antibodies directed against a NOVX protein can be used diagnostically to monitor protein levels in tissue as part of a clinical testing procedure, *e.g.*, to, for example, determine the efficacy of a given treatment regimen.

25 Detection can be facilitated by coupling (*i.e.*, physically linking) the antibody to a detectable substance. Examples of detectable substances include various enzymes, prosthetic groups, fluorescent materials, luminescent materials, bioluminescent materials, and radioactive materials. Examples of suitable enzymes include horseradish peroxidase, alkaline phosphatase, β-galactosidase, or acetylcholinesterase; examples of suitable

30 prosthetic group complexes include streptavidin/biotin and avidin/biotin; examples of suitable fluorescent materials include umbelliferone, fluorescein, fluorescein isothiocyanate, rhodamine, dichlorotriazinylamine fluorescein, dansyl chloride or phycoerythrin; an example of a luminescent material includes luminol; examples of

bioluminescent materials include luciferase, luciferin, and aequorin, and examples of suitable radioactive material include ^{125}I, ^{131}I, ^{35}S or ^{3}H.

Antibody Therapeutics

Antibodies of the invention, including polyclonal, monoclonal, humanized and fully human antibodies, may used as therapeutic agents. Such agents will generally be employed to treat or prevent a disease or pathology in a subject. An antibody preparation, preferably one having high specificity and high affinity for its target antigen, is administered to the subject and will generally have an effect due to its binding with the target. Such an effect may be one of two kinds, depending on the specific nature of the interaction between the given antibody molecule and the target antigen in question. In the first instance, administration of the antibody may abrogate or inhibit the binding of the target with an endogenous ligand to which it naturally binds. In this case, the antibody binds to the target and masks a binding site of the naturally occurring ligand, wherein the ligand serves as an effector molecule. Thus the receptor mediates a signal transduction pathway for which ligand is responsible.

Alternatively, the effect may be one in which the antibody elicits a physiological result by virtue of binding to an effector binding site on the target molecule. In this case the target, a receptor having an endogenous ligand which may be absent or defective in the disease or pathology, binds the antibody as a surrogate effector ligand, initiating a receptor-based signal transduction event by the receptor.

A therapeutically effective amount of an antibody of the invention relates generally to the amount needed to achieve a therapeutic objective. As noted above, this may be a binding interaction between the antibody and its target antigen that, in certain cases, interferes with the functioning of the target, and in other cases, promotes a physiological response. The amount required to be administered will furthermore depend on the binding affinity of the antibody for its specific antigen, and will also depend on the rate at which an administered antibody is depleted from the free volume other subject to which it is administered. Common ranges for therapeutically effective dosing of an antibody or antibody fragment of the invention may be, by way of nonlimiting example, from about 0.1 mg/kg body weight to about 50 mg/kg body weight. Common dosing frequencies may range, for example, from twice daily to once a week.

Pharmaceutical Compositions of Antibodies

Antibodies specifically binding a protein of the invention, as well as other molecules identified by the screening assays disclosed herein, can be administered for the treatment of various disorders in the form of pharmaceutical compositions. Principles and

5   considerations involved in preparing such compositions, as well as guidance in the choice of components are provided, for example, in Remington : The Science And Practice Of Pharmacy 19th ed. (Alfonso R. Gennaro, et al., editors) Mack Pub. Co., Easton, Pa. : 1995; Drug Absorption Enhancement : Concepts, Possibilities, Limitations, And Trends, Harwood Academic Publishers, Langhorne, Pa., 1994; and Peptide And Protein Drug

10   Delivery (Advances In Parenteral Sciences, Vol. 4), 1991, M. Dekker, New York.

If the antigenic protein is intracellular and whole antibodies are used as inhibitors, internalizing antibodies are preferred. However, liposomes can also be used to deliver the antibody, or an antibody fragment, into cells. Where antibody fragments are used, the smallest inhibitory fragment that specifically binds to the binding domain of the target

15   protein is preferred. For example, based upon the variable-region sequences of an antibody, peptide molecules can be designed that retain the ability to bind the target protein sequence. Such peptides can be synthesized chemically and/or produced by recombinant DNA technology. See, e.g., Marasco et al., Proc. Natl. Acad. Sci. USA, 90: 7889-7893 (1993). The formulation herein can also contain more than one active compound as

20   necessary for the particular indication being treated, preferably those with complementary activities that do not adversely affect each other. Alternatively, or in addition, the composition can comprise an agent that enhances its function, such as, for example, a cytotoxic agent, cytokine, chemotherapeutic agent, or growth-inhibitory agent. Such molecules are suitably present in combination in amounts that are effective for the purpose

25   intended.

The active ingredients can also be entrapped in microcapsules prepared, for example, by coacervation techniques or by interfacial polymerization, for example, hydroxymethylcellulose or gelatin-microcapsules and poly-(methylmethacrylate) microcapsules, respectively, in colloidal drug delivery systems (for example, liposomes,

30   albumin microspheres, microemulsions, nano-particles, and nanocapsules) or in macroemulsions.

The formulations to be used for in vivo administration must be sterile. This is readily accomplished by filtration through sterile filtration membranes.

Sustained-release preparations can be prepared. Suitable examples of sustained-release preparations include semipermeable matrices of solid hydrophobic polymers containing the antibody, which matrices are in the form of shaped articles, e.g., films, or microcapsules. Examples of sustained-release matrices include polyesters,

5   hydrogels (for example, poly(2-hydroxyethyl-methacrylate), or poly(vinylalcohol)), polylactides (U.S. Pat. No. 3,773,919), copolymers of L-glutamic acid and γ ethyl-L-glutamate, non-degradable ethylene-vinyl acetate, degradable lactic acid-glycolic acid copolymers such as the LUPRON DEPOT ™ (injectable microspheres composed of lactic acid-glycolic acid copolymer and leuprolide acetate), and

10   poly-D-(-)-3-hydroxybutyric acid. While polymers such as ethylene-vinyl acetate and lactic acid-glycolic acid enable release of molecules for over 100 days, certain hydrogels release proteins for shorter time periods.

ELISA Assay

An agent for detecting an analyte protein is an antibody capable of binding to an

15   analyte protein, preferably an antibody with a detectable label. Antibodies can be polyclonal, or more preferably, monoclonal. An intact antibody, or a fragment thereof (e.g., F_{ab} or $F_{(ab)2}$) can be used. The term "labeled", with regard to the probe or antibody, is intended to encompass direct labeling of the probe or antibody by coupling (i.e., physically linking) a detectable substance to the probe or antibody, as well as indirect labeling of the

20   probe or antibody by reactivity with another reagent that is directly labeled. Examples of indirect labeling include detection of a primary antibody using a fluorescently-labeled secondary antibody and end-labeling of a DNA probe with biotin such that it can be detected with fluorescently-labeled streptavidin. The term "biological sample" is intended to include tissues, cells and biological fluids isolated from a subject, as well as tissues, cells

25   and fluids present within a subject. Included within the usage of the term "biological sample", therefore, is blood and a fraction or component of blood including blood serum, blood plasma, or lymph. That is, the detection method of the invention can be used to detect an analyte mRNA, protein, or genomic DNA in a biological sample in vitro as well as in vivo. For example, in vitro techniques for detection of an analyte mRNA include

30   Northern hybridizations and in situ hybridizations. In vitro techniques for detection of an analyte protein include enzyme linked immunosorbent assays (ELISAs), Western blots, immunoprecipitations, and immunofluorescence. In vitro techniques for detection of an analyte genomic DNA include Southern hybridizations. Procedures for conducting

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immunoassays are described, for example in "ELISA: Theory and Practice: Methods in

Molecular Biology", Vol. 42, J. R. Crowther (Ed.) Human Press, Totowa, NJ, 1995;

"Immunoassay", E. Diamandis and T. Christopoulus, Academic Press, Inc., San Diego,

CA, 1996; and "Practice and Thory of Enzyme Immunoassays", P. Tijssen, Elsevier

5   Science Publishers, Amsterdam, 1985. Furthermore, *in vivo* techniques for detection of an

analyte protein include introducing into a subject a labeled anti-an analyte protein antibody.

For example, the antibody can be labeled with a radioactive marker whose presence and

location in a subject can be detected by standard imaging techniques.

NOVX Recombinant Expression Vectors and Host Cells

10   Another aspect of the invention pertains to vectors, preferably expression vectors,

containing a nucleic acid encoding a NOVX protein, or derivatives, fragments, analogs or

homologs thereof. As used herein, the term "vector" refers to a nucleic acid molecule

capable of transporting another nucleic acid to which it has been linked. One type of vector

is a "plasmid", which refers to a circular double stranded DNA loop into which additional

15   DNA segments can be ligated. Another type of vector is a viral vector, wherein additional

DNA segments can be ligated into the viral genome. Certain vectors are capable of

autonomous replication in a host cell into which they are introduced (*e.g.*, bacterial vectors

having a bacterial origin of replication and episomal mammalian vectors). Other vectors

(*e.g.*, non-episomal mammalian vectors) are integrated into the genome of a host cell upon

20   introduction into the host cell, and thereby are replicated along with the host genome.

Moreover, certain vectors are capable of directing the expression of genes to which they are

operatively-linked. Such vectors are referred to herein as "expression vectors". In general,

expression vectors of utility in recombinant DNA techniques are often in the form of

plasmids. In the present specification, "plasmid" and "vector" can be used interchangeably

25   as the plasmid is the most commonly used form of vector. However, the invention is

intended to include such other forms of expression vectors, such as viral vectors (*e.g.*,

replication defective retroviruses, adenoviruses and adeno-associated viruses), which serve

equivalent functions.

The recombinant expression vectors of the invention comprise a nucleic acid of the

30   invention in a form suitable for expression of the nucleic acid in a host cell, which means

that the recombinant expression vectors include one or more regulatory sequences, selected

on the basis of the host cells to be used for expression, that is operatively-linked to the

nucleic acid sequence to be expressed. Within a recombinant expression vector,

"operably-linked" is intended to mean that the nucleotide sequence of interest is linked to the regulatory sequence(s) in a manner that allows for expression of the nucleotide sequence (e.g., in an in vitro transcription/translation system or in a host cell when the vector is introduced into the host cell).

5   The term "regulatory sequence" is intended to includes promoters, enhancers and other expression control elements (e.g., polyadenylation signals). Such regulatory sequences are described, for example, in Goeddel, GENE EXPRESSION TECHNOLOGY: METHODS IN ENZYMOLOGY 185, Academic Press, San Diego, Calif. (1990). Regulatory sequences include those that direct constitutive expression of a nucleotide sequence in

10 many types of host cell and those that direct expression of the nucleotide sequence only in certain host cells (e.g., tissue-specific regulatory sequences). It will be appreciated by those skilled in the art that the design of the expression vector can depend on such factors as the choice of the host cell to be transformed, the level of expression of protein desired, etc. The expression vectors of the invention can be introduced into host cells to thereby

15 produce proteins or peptides, including fusion proteins or peptides, encoded by nucleic acids as described herein (e.g., NOVX proteins, mutant forms of NOVX proteins, fusion proteins, etc.).

   The recombinant expression vectors of the invention can be designed for expression of NOVX proteins in prokaryotic or eukaryotic cells. For example, NOVX proteins can be

20 expressed in bacterial cells such as Escherichia coli, insect cells (using baculovirus expression vectors) yeast cells or mammalian cells. Suitable host cells are discussed further in Goeddel, GENE EXPRESSION TECHNOLOGY: METHODS IN ENZYMOLOGY 185, Academic Press, San Diego, Calif. (1990). Alternatively, the recombinant expression vector can be transcribed and translated in vitro, for example using T7 promoter regulatory sequences

25 and T7 polymerase.

   Expression of proteins in prokaryotes is most often carried out in Escherichia coli with vectors containing constitutive or inducible promoters directing the expression of either fusion or non-fusion proteins. Fusion vectors add a number of amino acids to a protein encoded therein, usually to the amino terminus of the recombinant protein. Such

30 fusion vectors typically serve three purposes: (i) to increase expression of recombinant protein; (ii) to increase the solubility of the recombinant protein; and (iii) to aid in the purification of the recombinant protein by acting as a ligand in affinity purification. Often, in fusion expression vectors, a proteolytic cleavage site is introduced at the junction of the

fusion moiety and the recombinant protein to enable separation of the recombinant protein from the fusion moiety subsequent to purification of the fusion protein. Such enzymes, and their cognate recognition sequences, include Factor Xa, thrombin and enterokinase. Typical fusion expression vectors include pGEX (Pharmacia Biotech Inc; Smith and

5   Johnson, 1988. *Gene* 67: 31-40), pMAL (New England Biolabs, Beverly, Mass.) and pRIT5 (Pharmacia, Piscataway, N.J.) that fuse glutathione S-transferase (GST), maltose E binding protein, or protein A, respectively, to the target recombinant protein.

Examples of suitable inducible non-fusion *E. coli* expression vectors include pTrc (Amrann *et al.*, (1988) *Gene* 69:301-315) and pET 11d (Studier *et al.*, GENE EXPRESSION

10   TECHNOLOGY: METHODS IN ENZYMOLOGY 185, Academic Press, San Diego, Calif. (1990) 60-89).

One strategy to maximize recombinant protein expression in *E. coli* is to express the protein in a host bacteria with an impaired capacity to proteolytically cleave the recombinant protein. *See, e.g.*, Gottesman, GENE EXPRESSION TECHNOLOGY: METHODS IN

15   ENZYMOLOGY 185, Academic Press, San Diego, Calif. (1990) 119-128. Another strategy is to alter the nucleic acid sequence of the nucleic acid to be inserted into an expression vector so that the individual codons for each amino acid are those preferentially utilized in *E. coli* (*see, e.g.*, Wada, *et al.*, 1992. *Nucl. Acids Res.* 20: 2111-2118). Such alteration of nucleic acid sequences of the invention can be carried out by standard DNA synthesis techniques.

20   In another embodiment, the NOVX expression vector is a yeast expression vector. Examples of vectors for expression in yeast *Saccharomyces cerivisae* include pYepSec1 (Baldari, *et al.*, 1987. *EMBO J.* 6: 229-234), pMFa (Kurjan and Herskowitz, 1982. *Cell* 30: 933-943), pJRY88 (Schultz *et al.*, 1987. *Gene* 54: 113-123), pYES2 (Invitrogen Corporation, San Diego, Calif.), and picZ (InVitrogen Corp, San Diego, Calif.).

25   Alternatively, NOVX can be expressed in insect cells using baculovirus expression vectors. Baculovirus vectors available for expression of proteins in cultured insect cells (*e.g.*, SF9 cells) include the pAc series (Smith, *et al.*, 1983. *Mol. Cell. Biol.* 3: 2156-2165) and the pVL series (Lucklow and Summers, 1989. *Virology* 170: 31-39).

In yet another embodiment, a nucleic acid of the invention is expressed in

30   mammalian cells using a mammalian expression vector. Examples of mammalian expression vectors include pCDM8 (Seed, 1987. *Nature* 329: 840) and pMT2PC (Kaufman, *et al.*, 1987. *EMBO J.* 6: 187-195). When used in mammalian cells, the expression vector's control functions are often provided by viral regulatory elements. For example, commonly

used promoters are derived from polyoma, adenovirus 2, cytomegalovirus, and simian virus 40. For other suitable expression systems for both prokaryotic and eukaryotic cells see, *e.g.*, Chapters 16 and 17 of Sambrook, *et al.*, MOLECULAR CLONING: A LABORATORY MANUAL. 2nd ed., Cold Spring Harbor Laboratory, Cold Spring Harbor Laboratory Press,
5   Cold Spring Harbor, N.Y., 1989.

In another embodiment, the recombinant mammalian expression vector is capable of directing expression of the nucleic acid preferentially in a particular cell type (*e.g.*, tissue-specific regulatory elements are used to express the nucleic acid). Tissue-specific regulatory elements are known in the art. Non-limiting examples of suitable tissue-specific
10   promoters include the albumin promoter (liver-specific; Pinkert, *et al.*, 1987. *Genes Dev.* 1: 268-277), lymphoid-specific promoters (Calame and Eaton, 1988. *Adv. Immunol.* 43: 235-275), in particular promoters of T cell receptors (Winoto and Baltimore, 1989. *EMBO J.* 8: 729-733) and immunoglobulins (Banerji, *et al.*, 1983. *Cell* 33: 729-740; Queen and Baltimore, 1983. *Cell* 33: 741-748), neuron-specific promoters (*e.g.*, the neurofilament
15   promoter; Byrne and Ruddle, 1989. *Proc. Natl. Acad. Sci. USA* 86: 5473-5477), pancreas-specific promoters (Edlund, *et al.*, 1985. *Science* 230: 912-916), and mammary gland-specific promoters (*e.g.*, milk whey promoter; U.S. Pat. No. 4,873,316 and European Application Publication No. 264,166). Developmentally-regulated promoters are also encompassed, *e.g.*, the murine hox promoters (Kessel and Gruss, 1990. *Science* 249:
20   374-379) and the α-fetoprotein promoter (Campes and Tilghman, 1989. *Genes Dev.* 3: 537-546).

The invention further provides a recombinant expression vector comprising a DNA molecule of the invention cloned into the expression vector in an antisense orientation. That is, the DNA molecule is operatively-linked to a regulatory sequence in a manner that
25   allows for expression (by transcription of the DNA molecule) of an RNA molecule that is antisense to NOVX mRNA. Regulatory sequences operatively linked to a nucleic acid cloned in the antisense orientation can be chosen that direct the continuous expression of the antisense RNA molecule in a variety of cell types, for instance viral promoters and/or enhancers, or regulatory sequences can be chosen that direct constitutive, tissue specific or
30   cell type specific expression of antisense RNA. The antisense expression vector can be in the form of a recombinant plasmid, phagemid or attenuated virus in which antisense nucleic acids are produced under the control of a high efficiency regulatory region, the activity of which can be determined by the cell type into which the vector is introduced. For a

discussion of the regulation of gene expression using antisense genes *see, e.g.,* Weintraub, *et al.,* "Antisense RNA as a molecular tool for genetic analysis," *Reviews-Trends in Genetics,* Vol. 1(1) 1986.

Another aspect of the invention pertains to host cells into which a recombinant
expression vector of the invention has been introduced. The terms "host cell" and "recombinant host cell" are used interchangeably herein. It is understood that such terms refer not only to the particular subject cell but also to the progeny or potential progeny of such a cell. Because certain modifications may occur in succeeding generations due to either mutation or environmental influences, such progeny may not, in fact, be identical to the parent cell, but are still included within the scope of the term as used herein.

A host cell can be any prokaryotic or eukaryotic cell. For example, NOVX protein can be expressed in bacterial cells such as *E. coli,* insect cells, yeast or mammalian cells (such as Chinese hamster ovary cells (CHO) or COS cells). Other suitable host cells are known to those skilled in the art.

Vector DNA can be introduced into prokaryotic or eukaryotic cells via conventional transformation or transfection techniques. As used herein, the terms "transformation" and "transfection" are intended to refer to a variety of art-recognized techniques for introducing foreign nucleic acid (*e.g.,* DNA) into a host cell, including calcium phosphate or calcium chloride co-precipitation, DEAE-dextran-mediated transfection, lipofection, or electroporation. Suitable methods for transforming or transfecting host cells can be found in Sambrook, *et al.* (MOLECULAR CLONING: A LABORATORY MANUAL. 2nd ed., Cold Spring Harbor Laboratory, Cold Spring Harbor Laboratory Press, Cold Spring Harbor, N.Y., 1989), and other laboratory manuals.

For stable transfection of mammalian cells, it is known that, depending upon the expression vector and transfection technique used, only a small fraction of cells may integrate the foreign DNA into their genome. In order to identify and select these integrants, a gene that encodes a selectable marker (*e.g.,* resistance to antibiotics) is generally introduced into the host cells along with the gene of interest. Various selectable markers include those that confer resistance to drugs, such as G418, hygromycin and methotrexate. Nucleic acid encoding a selectable marker can be introduced into a host cell on the same vector as that encoding NOVX or can be introduced on a separate vector. Cells stably transfected with the introduced nucleic acid can be identified by drug selection

(*e.g.*, cells that have incorporated the selectable marker gene will survive, while the other cells die).

A host cell of the invention, such as a prokaryotic or eukaryotic host cell in culture, can be used to produce (*i.e.*, express) NOVX protein. Accordingly, the invention further

5   provides methods for producing NOVX protein using the host cells of the invention. In one embodiment, the method comprises culturing the host cell of invention (into which a recombinant expression vector encoding NOVX protein has been introduced) in a suitable medium such that NOVX protein is produced. In another embodiment, the method further comprises isolating NOVX protein from the medium or the host cell.

10   **Transgenic NOVX Animals**

The host cells of the invention can also be used to produce non-human transgenic animals. For example, in one embodiment, a host cell of the invention is a fertilized oocyte or an embryonic stem cell into which NOVX protein-coding sequences have been introduced. Such host cells can then be used to create non-human transgenic animals in

15   which exogenous NOVX sequences have been introduced into their genome or homologous recombinant animals in which endogenous NOVX sequences have been altered. Such animals are useful for studying the function and/or activity of NOVX protein and for identifying and/or evaluating modulators of NOVX protein activity. As used herein, a "transgenic animal" is a non-human animal, preferably a mammal, more

20   preferably a rodent such as a rat or mouse, in which one or more of the cells of the animal includes a transgene. Other examples of transgenic animals include non-human primates, sheep, dogs, cows, goats, chickens, amphibians, *etc.* A transgene is exogenous DNA that is integrated into the genome of a cell from which a transgenic animal develops and that remains in the genome of the mature animal, thereby directing the expression of an

25   encoded gene product in one or more cell types or tissues of the transgenic animal. As used herein, a "homologous recombinant animal" is a non-human animal, preferably a mammal, more preferably a mouse, in which an endogenous NOVX gene has been altered by homologous recombination between the endogenous gene and an exogenous DNA molecule introduced into a cell of the animal, *e.g.*, an embryonic cell of the animal, prior to

30   development of the animal.

A transgenic animal of the invention can be created by introducing NOVX-encoding nucleic acid into the male pronuclei of a fertilized oocyte (*e.g.*, by microinjection, retroviral infection) and allowing the oocyte to develop in a pseudopregnant

female foster animal. The human NOVX cDNA sequences, i.e., any one of SEQ ID NO:2n-1, wherein n is an integer between 1 and 124, can be introduced as a transgene into the genome of a non-human animal. Alternatively, a non-human homologue of the human NOVX gene, such as a mouse NOVX gene, can be isolated based on hybridization to the

5   human NOVX cDNA (described further *supra*) and used as a transgene. Intronic sequences and polyadenylation signals can also be included in the transgene to increase the efficiency of expression of the transgene. A tissue-specific regulatory sequence(s) can be operably-linked to the NOVX transgene to direct expression of NOVX protein to particular cells. Methods for generating transgenic animals via embryo manipulation and

10   microinjection, particularly animals such as mice, have become conventional in the art and are described, for example, in U.S. Patent Nos. 4,736,866; 4,870,009; and 4,873,191; and Hogan, 1986. In: MANIPULATING THE MOUSE EMBRYO, Cold Spring Harbor Laboratory Press, Cold Spring Harbor, N.Y. Similar methods are used for production of other transgenic animals. A transgenic founder animal can be identified based upon the presence

15   of the NOVX transgene in its genome and/or expression of NOVX mRNA in tissues or cells of the animals. A transgenic founder animal can then be used to breed additional animals carrying the transgene. Moreover, transgenic animals carrying a transgene-encoding NOVX protein can further be bred to other transgenic animals carrying other transgenes.

20   To create a homologous recombinant animal, a vector is prepared which contains at least a portion of a NOVX gene into which a deletion, addition or substitution has been introduced to thereby alter, *e.g.*, functionally disrupt, the NOVX gene. The NOVX gene can be a human gene (*e.g.*, the cDNA of any one of SEQ ID NO:2n-1, wherein n is an integer between 1 and 124), but more preferably, is a non-human homologue of a human

25   NOVX gene. For example, a mouse homologue of human NOVX gene of SEQ ID NO:2n-1, wherein n is an integer between 1 and 124, can be used to construct a homologous recombination vector suitable for altering an endogenous NOVX gene in the mouse genome. In one embodiment, the vector is designed such that, upon homologous recombination, the endogenous NOVX gene is functionally disrupted (*i.e.*, no longer

30   encodes a functional protein; also referred to as a "knock out" vector).

Alternatively, the vector can be designed such that, upon homologous recombination, the endogenous NOVX gene is mutated or otherwise altered but still encodes functional protein (*e.g.*, the upstream regulatory region can be altered to thereby

alter the expression of the endogenous NOVX protein). In the homologous recombination vector, the altered portion of the NOVX gene is flanked at its 5'- and 3'-termini by additional nucleic acid of the NOVX gene to allow for homologous recombination to occur between the exogenous NOVX gene carried by the vector and an endogenous NOVX gene

5   in an embryonic stem cell. The additional flanking NOVX nucleic acid is of sufficient length for successful homologous recombination with the endogenous gene. Typically, several kilobases of flanking DNA (both at the 5'- and 3'-termini) are included in the vector. *See, e.g.,* Thomas, *et al.,* 1987. *Cell* 51: 503 for a description of homologous recombination vectors. The vector is ten introduced into an embryonic stem cell line (*e.g.,*

10   by electroporation) and cells in which the introduced NOVX gene has homologously-recombined with the endogenous NOVX gene are selected. *See, e.g.,* Li, *et al.,* 1992. *Cell* 69: 915.

The selected cells are then injected into a blastocyst of an animal (*e.g.,* a mouse) to form aggregation chimeras. *See, e.g.,* Bradley, 1987. In: TERATOCARCINOMAS AND

15   EMBRYONIC STEM CELLS: A PRACTICAL APPROACH, Robertson, ed. IRL, Oxford, pp. 113-152. A chimeric embryo can then be implanted into a suitable pseudopregnant female foster animal and the embryo brought to term. Progeny harboring the homologously-recombined DNA in their germ cells can be used to breed animals in which all cells of the animal contain the homologously-recombined DNA by germline

20   transmission of the transgene. Methods for constructing homologous recombination vectors and homologous recombinant animals are described further in Bradley, 1991. *Curr. Opin. Biotechnol.* 2: 823-829; PCT International Publication Nos.: WO 90/11354; WO 91/01140; WO 92/0968; and WO 93/04169.

In another embodiment, transgenic non-humans animals can be produced that

25   contain selected systems that allow for regulated expression of the transgene. One example of such a system is the cre/loxP recombinase system of bacteriophage P1. For a description of the cre/loxP recombinase system, *See, e.g.,* Lakso, *et al.,* 1992. *Proc. Natl. Acad. Sci. USA* 89: 6232-6236. Another example of a recombinase system is the FLP recombinase system of *Saccharomyces cerevisiae. See,* O'Gorman, *et al.,* 1991. *Science* 251:1351-1355.

30   If a cre/loxP recombinase system is used to regulate expression of the transgene, animals containing transgenes encoding both the Cre recombinase and a selected protein are required. Such animals can be provided through the construction of "double" transgenic

animals, *e.g.*, by mating two transgenic animals, one containing a transgene encoding a selected protein and the other containing a transgene encoding a recombinase.

Clones of the non-human transgenic animals described herein can also be produced according to the methods described in Wilmut, *et al.*, 1997. *Nature* 385: 810-813. In brief,
5   a cell (*e.g.*, a somatic cell) from the transgenic animal can be isolated and induced to exit the growth cycle and enter G_0 phase. The quiescent cell can then be fused, *e.g.*, through the use of electrical pulses, to an enucleated oocyte from an animal of the same species from which the quiescent cell is isolated. The reconstructed oocyte is then cultured such that it develops to morula or blastocyte and then transferred to pseudopregnant female foster
10   animal. The offspring borne of this female foster animal will be a clone of the animal from which the cell (*e.g.*, the somatic cell) is isolated.

Pharmaceutical Compositions

The NOVX nucleic acid molecules, NOVX proteins, and anti-NOVX antibodies (also referred to herein as "active compounds") of the invention, and derivatives, fragments,
15   analogs and homologs thereof, can be incorporated into pharmaceutical compositions suitable for administration. Such compositions typically comprise the nucleic acid molecule, protein, or antibody and a pharmaceutically acceptable carrier. As used herein, "pharmaceutically acceptable carrier" is intended to include any and all solvents, dispersion media, coatings, antibacterial and antifungal agents, isotonic and absorption delaying
20   agents, and the like, compatible with pharmaceutical administration. Suitable carriers are described in the most recent edition of Remington's Pharmaceutical Sciences, a standard reference text in the field, which is incorporated herein by reference. Preferred examples of such carriers or diluents include, but are not limited to, water, saline, finger's solutions, dextrose solution, and 5% human serum albumin. Liposomes and non-aqueous vehicles
25   such as fixed oils may also be used. The use of such media and agents for pharmaceutically active substances is well known in the art. Except insofar as any conventional media or agent is incompatible with the active compound, use thereof in the compositions is contemplated. Supplementary active compounds can also be incorporated into the compositions.

30   A pharmaceutical composition of the invention is formulated to be compatible with its intended route of administration. Examples of routes of administration include parenteral, *e.g.*, intravenous, intradermal, subcutaneous, oral (*e.g.*, inhalation), transdermal (*i.e.*, topical), transmucosal, and rectal administration. Solutions or suspensions used for

parenteral, intradermal, or subcutaneous application can include the following components:
a sterile diluent such as water for injection, saline solution, fixed oils, polyethylene glycols,
glycerine, propylene glycol or other synthetic solvents; antibacterial agents such as benzyl
alcohol or methyl parabens; antioxidants such as ascorbic acid or sodium bisulfite;

5　　chelating agents such as ethylenediaminetetraacetic acid (EDTA); buffers such as acetates,
citrates or phosphates, and agents for the adjustment of tonicity such as sodium chloride or
dextrose. The pH can be adjusted with acids or bases, such as hydrochloric acid or sodium
hydroxide. The parenteral preparation can be enclosed in ampoules, disposable syringes or
multiple dose vials made of glass or plastic.

10　　　　　Pharmaceutical compositions suitable for injectable use include sterile aqueous
solutions (where water soluble) or dispersions and sterile powders for the extemporaneous
preparation of sterile injectable solutions or dispersion. For intravenous administration,
suitable carriers include physiological saline, bacteriostatic water, Cremophor EL™ (BASF,
Parsippany, N.J.) or phosphate buffered saline (PBS). In all cases, the composition must be

15　　sterile and should be fluid to the extent that easy syringeability exists. It must be stable
under the conditions of manufacture and storage and must be preserved against the
contaminating action of microorganisms such as bacteria and fungi. The carrier can be a
solvent or dispersion medium containing, for example, water, ethanol, polyol (for example,
glycerol, propylene glycol, and liquid polyethylene glycol, and the like), and suitable

20　　mixtures thereof. The proper fluidity can be maintained, for example, by the use of a
coating such as lecithin, by the maintenance of the required particle size in the case of
dispersion and by the use of surfactants. Prevention of the action of microorganisms can be
achieved by various antibacterial and antifungal agents, for example, parabens,
chlorobutanol, phenol, ascorbic acid, thimerosal, and the like. In many cases, it will be

25　　preferable to include isotonic agents, for example, sugars, polyalcohols such as manitol,
sorbitol, sodium chloride in the composition. Prolonged absorption of the injectable
compositions can be brought about by including in the composition an agent which delays
absorption, for example, aluminum monostearate and gelatin.

　　　　　Sterile injectable solutions can be prepared by incorporating the active compound
30　　(e.g., a NOVX protein or anti-NOVX antibody) in the required amount in an appropriate
solvent with one or a combination of ingredients enumerated above, as required, followed
by filtered sterilization. Generally, dispersions are prepared by incorporating the active
compound into a sterile vehicle that contains a basic dispersion medium and the required

other ingredients from those enumerated above. In the case of sterile powders for the
preparation of sterile injectable solutions, methods of preparation are vacuum drying and
freeze-drying that yields a powder of the active ingredient plus any additional desired
ingredient from a previously sterile-filtered solution thereof.

5   Oral compositions generally include an inert diluent or an edible carrier. They can
be enclosed in gelatin capsules or compressed into tablets. For the purpose of oral
therapeutic administration, the active compound can be incorporated with excipients and
used in the form of tablets, troches, or capsules. Oral compositions can also be prepared
using a fluid carrier for use as a mouthwash, wherein the compound in the fluid carrier is
10   applied orally and swished and expectorated or swallowed. Pharmaceutically compatible
binding agents, and/or adjuvant materials can be included as part of the composition. The
tablets, pills, capsules, troches and the like can contain any of the following ingredients, or
compounds of a similar nature: a binder such as microcrystalline cellulose, gum tragacanth
or gelatin; an excipient such as starch or lactose, a disintegrating agent such as alginic acid,
15   Primogel, or corn starch; a lubricant such as magnesium stearate or Sterotes; a glidant such
as colloidal silicon dioxide; a sweetening agent such as sucrose or saccharin; or a flavoring
agent such as peppermint, methyl salicylate, or orange flavoring.

          For administration by inhalation, the compounds are delivered in the form of an
aerosol spray from pressured container or dispenser which contains a suitable propellant,
20   e.g., a gas such as carbon dioxide, or a nebulizer.

          Systemic administration can also be by transmucosal or transdermal means. For
transmucosal or transdermal administration, penetrants appropriate to the barrier to be
permeated are used in the formulation. Such penetrants are generally known in the art, and
include, for example, for transmucosal administration, detergents, bile salts, and fusidic
25   acid derivatives. Transmucosal administration can be accomplished through the use of
nasal sprays or suppositories. For transdermal administration, the active compounds are
formulated into ointments, salves, gels, or creams as generally known in the art.

          The compounds can also be prepared in the form of suppositories (e.g., with
conventional suppository bases such as cocoa butter and other glycerides) or retention
30   enemas for rectal delivery.

          In one embodiment, the active compounds are prepared with carriers that will
protect the compound against rapid elimination from the body, such as a controlled release
formulation, including implants and microencapsulated delivery systems. Biodegradable,

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biocompatible polymers can be used, such as ethylene vinyl acetate, polyanhydrides, polyglycolic acid, collagen, polyorthoesters, and polylactic acid. Methods for preparation of such formulations will be apparent to those skilled in the art. The materials can also be obtained commercially from Alza Corporation and Nova Pharmaceuticals, Inc. Liposomal

5   suspensions (including liposomes targeted to infected cells with monoclonal antibodies to viral antigens) can also be used as pharmaceutically acceptable carriers. These can be prepared according to methods known to those skilled in the art, for example, as described in U.S. Patent No. 4,522,811.

It is especially advantageous to formulate oral or parenteral compositions in dosage
10   unit form for ease of administration and uniformity of dosage. Dosage unit form as used herein refers to physically discrete units suited as unitary dosages for the subject to be treated; each unit containing a predetermined quantity of active compound calculated to produce the desired therapeutic effect in association with the required pharmaceutical carrier. The specification for the dosage unit forms of the invention are dictated by and
15   directly dependent on the unique characteristics of the active compound and the particular therapeutic effect to be achieved, and the limitations inherent in the art of compounding such an active compound for the treatment of individuals.

The nucleic acid molecules of the invention can be inserted into vectors and used as gene therapy vectors. Gene therapy vectors can be delivered to a subject by, for example,
20   intravenous injection, local administration (see, e.g., U.S. Patent No. 5,328,470) or by stereotactic injection (see, e.g., Chen, et al., 1994. Proc. Natl. Acad. Sci. USA 91: 3054-3057). The pharmaceutical preparation of the gene therapy vector can include the gene therapy vector in an acceptable diluent, or can comprise a slow release matrix in which the gene delivery vehicle is imbedded. Alternatively, where the complete gene
25   delivery vector can be produced intact from recombinant cells, e.g., retroviral vectors, the pharmaceutical preparation can include one or more cells that produce the gene delivery system.

The pharmaceutical compositions can be included in a container, pack, or dispenser together with instructions for administration.

30   **Screening and Detection Methods**

The isolated nucleic acid molecules of the invention can be used to express NOVX protein (e.g., via a recombinant expression vector in a host cell in gene therapy applications), to detect NOVX mRNA (e.g., in a biological sample) or a genetic lesion in a

NOVX gene, and to modulate NOVX activity, as described further, below. In addition, the NOVX proteins can be used to screen drugs or compounds that modulate the NOVX protein activity or expression as well as to treat disorders characterized by insufficient or excessive production of NOVX protein or production of NOVX protein forms that have

5   decreased or aberrant activity compared to NOVX wild-type protein (*e.g.*; diabetes (regulates insulin release); obesity (binds and transport lipids); metabolic disturbances associated with obesity, the metabolic syndrome X as well as anorexia and wasting disorders associated with chronic diseases and various cancers, and infectious disease(possesses anti-microbial activity) and the various dyslipidemias. In addition, the

10   anti-NOVX antibodies of the invention can be used to detect and isolate NOVX proteins and modulate NOVX activity. In yet a further aspect, the invention can be used in methods to influence appetite, absorption of nutrients and the disposition of metabolic substrates in both a positive and negative fashion.

The invention further pertains to novel agents identified by the screening assays

15   described herein and uses thereof for treatments as described, *supra.*

Screening Assays

The invention provides a method (also referred to herein as a "screening assay") for identifying modulators, *i.e.*, candidate or test compounds or agents (*e.g.*, peptides, peptidomimetics, small molecules or other drugs) that bind to NOVX proteins or have a

20   stimulatory or inhibitory effect on, *e.g.*, NOVX protein expression or NOVX protein activity. The invention also includes compounds identified in the screening assays described herein.

In one embodiment, the invention provides assays for screening candidate or test compounds which bind to or modulate the activity of the membrane-bound form of a

25   NOVX protein or polypeptide or biologically-active portion thereof. The test compounds of the invention can be obtained using any of the numerous approaches in combinatorial library methods known in the art, including: biological libraries; spatially addressable parallel solid phase or solution phase libraries; synthetic library methods requiring deconvolution; the "one-bead one-compound" library method; and synthetic library

30   methods using affinity chromatography selection. The biological library approach is limited to peptide libraries, while the other four approaches are applicable to peptide, non-peptide oligomer or small molecule libraries of compounds. *See, e.g.,* Lam, 1997. *Anticancer Drug Design* 12: 145.

A "small molecule" as used herein, is meant to refer to a composition that has a molecular weight of less than about 5 kD and most preferably less than about 4 kD. Small molecules can be, *e.g.*, nucleic acids, peptides, polypeptides, peptidomimetics, carbohydrates, lipids or other organic or inorganic molecules. Libraries of chemical and/or

5   biological mixtures, such as fungal, bacterial, or algal extracts, are known in the art and can be screened with any of the assays of the invention.

Examples of methods for the synthesis of molecular libraries can be found in the art, for example in: DeWitt, *et al.*, 1993. *Proc. Natl. Acad. Sci. U.S.A.* 90: 6909; Erb, *et al.*, 1994. *Proc. Natl. Acad. Sci. U.S.A.* 91: 11422; Zuckermann, *et al.*, 1994. *J. Med. Chem.* 37:

10   2678; Cho, *et al.*, 1993. *Science* 261: 1303; Carrell, *et al.*, 1994. *Angew. Chem. Int. Ed. Engl.* 33: 2059; Carell, *et al.*, 1994. *Angew. Chem. Int. Ed. Engl.* 33: 2061; and Gallop, *et al.*, 1994. *J. Med. Chem.* 37: 1233.

Libraries of compounds may be presented in solution (*e.g.*, Houghten, 1992. *Biotechniques* 13: 412-421), or on beads (Lam, 1991. *Nature* 354: 82-84), on chips (Fodor,

15   1993. *Nature* 364: 555-556), bacteria (Ladner, U.S. Patent No. 5,223,409), spores (Ladner, U.S. Patent 5,233,409), plasmids (Cull, *et al.*, 1992. *Proc. Natl. Acad. Sci. USA* 89: 1865-1869) or on phage (Scott and Smith, 1990. *Science* 249: 386-390; Devlin, 1990. *Science* 249: 404-406; Cwirla, *et al.*, 1990. *Proc. Natl. Acad. Sci. U.S.A.* 87: 6378-6382; Felici, 1991. *J. Mol. Biol.* 222: 301-310; Ladner, U.S. Patent No. 5,233,409.).

20   In one embodiment, an assay is a cell-based assay in which a cell which expresses a membrane-bound form of NOVX protein, or a biologically-active portion thereof, on the cell surface is contacted with a test compound and the ability of the test compound to bind to a NOVX protein determined. The cell, for example, can of mammalian origin or a yeast cell. Determining the ability of the test compound to bind to the NOVX protein can be

25   accomplished, for example, by coupling the test compound with a radioisotope or enzymatic label such that binding of the test compound to the NOVX protein or biologically-active portion thereof can be determined by detecting the labeled compound in a complex. For example, test compounds can be labeled with ^{125}I, ^{35}S, ^{14}C, or ^{3}H, either directly or indirectly, and the radioisotope detected by direct counting of radioemission or

30   by scintillation counting. Alternatively, test compounds can be enzymatically-labeled with, for example, horseradish peroxidase, alkaline phosphatase, or luciferase, and the enzymatic label detected by determination of conversion of an appropriate substrate to product. In one embodiment, the assay comprises contacting a cell which expresses a membrane-bound

form of NOVX protein, or a biologically-active portion thereof, on the cell surface with a
known compound which binds NOVX to form an assay mixture, contacting the assay
mixture with a test compound, and determining the ability of the test compound to interact
with a NOVX protein, wherein determining the ability of the test compound to interact with
5   a NOVX protein comprises determining the ability of the test compound to preferentially
bind to NOVX protein or a biologically-active portion thereof as compared to the known
compound.

In another embodiment, an assay is a cell-based assay comprising contacting a cell
expressing a membrane-bound form of NOVX protein, or a biologically-active portion
10 thereof, on the cell surface with a test compound and determining the ability of the test
compound to modulate (e.g., stimulate or inhibit) the activity of the NOVX protein or
biologically-active portion thereof. Determining the ability of the test compound to
modulate the activity of NOVX or a biologically-active portion thereof can be
accomplished, for example, by determining the ability of the NOVX protein to bind to or
15 interact with a NOVX target molecule. As used herein, a "target molecule" is a molecule
with which a NOVX protein binds or interacts in nature, for example, a molecule on the
surface of a cell which expresses a NOVX interacting protein, a molecule on the surface of
a second cell, a molecule in the extracellular milieu, a molecule associated with the internal
surface of a cell membrane or a cytoplasmic molecule. A NOVX target molecule can be a
20 non-NOVX molecule or a NOVX protein or polypeptide of the invention. In one
embodiment, a NOVX target molecule is a component of a signal transduction pathway
that facilitates transduction of an extracellular signal (e.g. a signal generated by binding of
a compound to a membrane-bound NOVX molecule) through the cell membrane and into
the cell. The target, for example, can be a second intercellular protein that has catalytic
25 activity or a protein that facilitates the association of downstream signaling molecules with
NOVX.

Determining the ability of the NOVX protein to bind to or interact with a NOVX
target molecule can be accomplished by one of the methods described above for
determining direct binding. In one embodiment, determining the ability of the NOVX
30 protein to bind to or interact with a NOVX target molecule can be accomplished by
determining the activity of the target molecule. For example, the activity of the target
molecule can be determined by detecting induction of a cellular second messenger of the
target (i.e. intracellular Ca^{2+}, diacylglycerol, IP_3, etc.), detecting catalytic/enzymatic

activity of the target an appropriate substrate, detecting the induction of a reporter gene (comprising a NOVX-responsive regulatory element operatively linked to a nucleic acid encoding a detectable marker, *e.g.*, luciferase), or detecting a cellular response, for example, cell survival, cellular differentiation, or cell proliferation.

5   In yet another embodiment, an assay of the invention is a cell-free assay comprising contacting a NOVX protein or biologically-active portion thereof with a test compound and determining the ability of the test compound to bind to the NOVX protein or biologically-active portion thereof. Binding of the test compound to the NOVX protein can be determined either directly or indirectly as described above. In one such embodiment,

10   the assay comprises contacting the NOVX protein or biologically-active portion thereof with a known compound which binds NOVX to form an assay mixture, contacting the assay mixture with a test compound, and determining the ability of the test compound to interact with a NOVX protein, wherein determining the ability of the test compound to interact with a NOVX protein comprises determining the ability of the test compound to

15   preferentially bind to NOVX or biologically-active portion thereof as compared to the known compound.

In still another embodiment, an assay is a cell-free assay comprising contacting NOVX protein or biologically-active portion thereof with a test compound and determining the ability of the test compound to modulate (*e.g.* stimulate or inhibit) the activity of the

20   NOVX protein or biologically-active portion thereof. Determining the ability of the test compound to modulate the activity of NOVX can be accomplished, for example, by determining the ability of the NOVX protein to bind to a NOVX target molecule by one of the methods described above for determining direct binding. In an alternative embodiment, determining the ability of the test compound to modulate the activity of NOVX protein can

25   be accomplished by determining the ability of the NOVX protein further modulate a NOVX target molecule. For example, the catalytic/enzymatic activity of the target molecule on an appropriate substrate can be determined as described, *supra*.

In yet another embodiment, the cell-free assay comprises contacting the NOVX protein or biologically-active portion thereof with a known compound which binds NOVX

30   protein to form an assay mixture, contacting the assay mixture with a test compound, and determining the ability of the test compound to interact with a NOVX protein, wherein determining the ability of the test compound to interact with a NOVX protein comprises

determining the ability of the NOVX protein to preferentially bind to or modulate the activity of a NOVX target molecule.

The cell-free assays of the invention are amenable to use of both the soluble form or the membrane-bound form of NOVX protein. In the case of cell-free assays comprising the

5   membrane-bound form of NOVX protein, it may be desirable to utilize a solubilizing agent such that the membrane-bound form of NOVX protein is maintained in solution. Examples of such solubilizing agents include non-ionic detergents such as n-octylglucoside, n-dodecylglucoside, n-dodecylmaltoside, octanoyl-N-methylglucamide, decanoyl-N-methylglucamide, Triton® X-100, Triton® X-114, Thesit®,

10   Isotridecypoly(ethylene glycol ether)$_n$, N-dodecyl--N,N-dimethyl-3-ammonio-1-propane sulfonate, 3-(3-cholamidopropyl) dimethylammminiol-1-propane sulfonate (CHAPS), or 3-(3-cholamidopropyl)dimethylammminiol-2-hydroxy-1-propane sulfonate (CHAPSO).

In more than one embodiment of the above assay methods of the invention, it may be desirable to immobilize either NOVX protein or its target molecule to facilitate

15   separation of complexed from uncomplexed forms of one or both of the proteins, as well as to accommodate automation of the assay. Binding of a test compound to NOVX protein, or interaction of NOVX protein with a target molecule in the presence and absence of a candidate compound, can be accomplished in any vessel suitable for containing the reactants. Examples of such vessels include microtiter plates, test tubes, and

20   micro-centrifuge tubes. In one embodiment, a fusion protein can be provided that adds a domain that allows one or both of the proteins to be bound to a matrix. For example, GST-NOVX fusion proteins or GST-target fusion proteins can be adsorbed onto glutathione sepharose beads (Sigma Chemical, St. Louis, MO) or glutathione derivatized microtiter plates, that are then combined with the test compound or the test compound and

25   either the non-adsorbed target protein or NOVX protein, and the mixture is incubated under conditions conducive to complex formation (e.g., at physiological conditions for salt and pH). Following incubation, the beads or microtiter plate wells are washed to remove any unbound components, the matrix immobilized in the case of beads, complex determined either directly or indirectly, for example, as described, supra. Alternatively, the complexes

30   can be dissociated from the matrix, and the level of NOVX protein binding or activity determined using standard techniques.

Other techniques for immobilizing proteins on matrices can also be used in the screening assays of the invention. For example, either the NOVX protein or its target

molecule can be immobilized utilizing conjugation of biotin and streptavidin. Biotinylated NOVX protein or target molecules can be prepared from biotin-NHS (N-hydroxy-succinimide) using techniques well-known within the art (e.g., biotinylation kit, Pierce Chemicals, Rockford, Ill.), and immobilized in the wells of streptavidin-coated

5　　96 well plates (Pierce Chemical). Alternatively, antibodies reactive with NOVX protein or target molecules, but which do not interfere with binding of the NOVX protein to its target molecule, can be derivatized to the wells of the plate, and unbound target or NOVX protein trapped in the wells by antibody conjugation. Methods for detecting such complexes, in addition to those described above for the GST-immobilized complexes, include

10　　immunodetection of complexes using antibodies reactive with the NOVX protein or target molecule, as well as enzyme-linked assays that rely on detecting an enzymatic activity associated with the NOVX protein or target molecule.

　　　　In another embodiment, modulators of NOVX protein expression are identified in a method wherein a cell is contacted with a candidate compound and the expression of

15　　NOVX mRNA or protein in the cell is determined. The level of expression of NOVX mRNA or protein in the presence of the candidate compound is compared to the level of expression of NOVX mRNA or protein in the absence of the candidate compound. The candidate compound can then be identified as a modulator of NOVX mRNA or protein expression based upon this comparison. For example, when expression of NOVX mRNA

20　　or protein is greater (i.e., statistically significantly greater) in the presence of the candidate compound than in its absence, the candidate compound is identified as a stimulator of NOVX mRNA or protein expression. Alternatively, when expression of NOVX mRNA or protein is less (statistically significantly less) in the presence of the candidate compound than in its absence, the candidate compound is identified as an inhibitor of NOVX mRNA

25　　or protein expression. The level of NOVX mRNA or protein expression in the cells can be determined by methods described herein for detecting NOVX mRNA or protein.

　　　　In yet another aspect of the invention, the NOVX proteins can be used as "bait proteins" in a two-hybrid assay or three hybrid assay (see, e.g., U.S. Patent No. 5,283,317; Zervos, et al., 1993. Cell 72: 223-232; Madura, et al., 1993. J. Biol. Chem. 268:

30　　12046-12054; Bartel, et al., 1993. Biotechniques 14: 920-924; Iwabuchi, et al., 1993. Oncogene 8: 1693-1696; and Brent WO 94/10300), to identify other proteins that bind to or interact with NOVX ("NOVX-binding proteins" or "NOVX-bp") and modulate NOVX activity. Such NOVX-binding proteins are also involved in the propagation of signals by

the NOVX proteins as, for example, upstream or downstream elements of the NOVX pathway.

The two-hybrid system is based on the modular nature of most transcription factors, which consist of separable DNA-binding and activation domains. Briefly, the assay utilizes

5   two different DNA constructs. In one construct, the gene that codes for NOVX is fused to a gene encoding the DNA binding domain of a known transcription factor (*e.g.*, GAL-4). In the other construct, a DNA sequence, from a library of DNA sequences, that encodes an unidentified protein ("prey" or "sample") is fused to a gene that codes for the activation domain of the known transcription factor. If the "bait" and the "prey" proteins are able to

10   interact, *in vivo*, forming a NOVX-dependent complex, the DNA-binding and activation domains of the transcription factor are brought into close proximity. This proximity allows transcription of a reporter gene (*e.g.*, LacZ) that is operably linked to a transcriptional regulatory site responsive to the transcription factor. Expression of the reporter gene can be detected and cell colonies containing the functional transcription factor can be isolated

15   and used to obtain the cloned gene that encodes the protein which interacts with NOVX.

The invention further pertains to novel agents identified by the aforementioned screening assays and uses thereof for treatments as described herein.

Detection Assays

Portions or fragments of the cDNA sequences identified herein (and the

20   corresponding complete gene sequences) can be used in numerous ways as polynucleotide reagents. By way of example, and not of limitation, these sequences can be used to: (*i*) map their respective genes on a chromosome; and, thus, locate gene regions associated with genetic disease; (*ii*) identify an individual from a minute biological sample (tissue typing); and (*iii*) aid in forensic identification of a biological sample. Some of these applications

25   are described in the subsections, below.

Chromosome Mapping

Once the sequence (or a portion of the sequence) of a gene has been isolated, this sequence can be used to map the location of the gene on a chromosome. This process is called chromosome mapping. Accordingly, portions or fragments of the NOVX sequences

30   of SEQ ID NO:2*n*-1, wherein *n* is an integer between 1 and 124, or fragments or derivatives thereof, can be used to map the location of the NOVX genes, respectively, on a

chromosome. The mapping of the NOVX sequences to chromosomes is an important first step in correlating these sequences with genes associated with disease.

Briefly, NOVX genes can be mapped to chromosomes by preparing PCR primers (preferably 15-25 bp in length) from the NOVX sequences. Computer analysis of the

5   NOVX, sequences can be used to rapidly select primers that do not span more than one exon in the genomic DNA, thus complicating the amplification process. These primers can then be used for PCR screening of somatic cell hybrids containing individual human chromosomes. Only those hybrids containing the human gene corresponding to the NOVX sequences will yield an amplified fragment.

10   Somatic cell hybrids are prepared by fusing somatic cells from different mammals (*e.g.*, human and mouse cells). As hybrids of human and mouse cells grow and divide, they gradually lose human chromosomes in random order, but retain the mouse chromosomes. By using media in which mouse cells cannot grow, because they lack a particular enzyme, but in which human cells can, the one human chromosome that contains the gene encoding

15   the needed enzyme will be retained. By using various media, panels of hybrid cell lines can be established. Each cell line in a panel contains either a single human chromosome or a small number of human chromosomes, and a full set of mouse chromosomes, allowing easy mapping of individual genes to specific human chromosomes. *See, e.g.*, D'Eustachio, *et al.*, 1983. *Science* 220: 919-924. Somatic cell hybrids containing only fragments of

20   human chromosomes can also be produced by using human chromosomes with translocations and deletions.

PCR mapping of somatic cell hybrids is a rapid procedure for assigning a particular sequence to a particular chromosome. Three or more sequences can be assigned per day using a single thermal cycler. Using the NOVX sequences to design oligonucleotide

25   primers, sub-localization can be achieved with panels of fragments from specific chromosomes.

Fluorescence *in situ* hybridization (FISH) of a DNA sequence to a metaphase chromosomal spread can further be used to provide a precise chromosomal location in one step. Chromosome spreads can be made using cells whose division has been blocked in

30   metaphase by a chemical like colcemid that disrupts the mitotic spindle. The chromosomes can be treated briefly with trypsin, and then stained with Giemsa. A pattern of light and dark bands develops on each chromosome, so that the chromosomes can be identified individually. The FISH technique can be used with a DNA sequence as short as 500 or 600

bases. However, clones larger than 1,000 bases have a higher likelihood of binding to a unique chromosomal location with sufficient signal intensity for simple detection. Preferably 1,000 bases, and more preferably 2,000 bases, will suffice to get good results at a reasonable amount of time. For a review of this technique, *see,* Verma, *et al.,* HUMAN

5   CHROMOSOMES: A MANUAL OF BASIC TECHNIQUES (Pergamon Press, New York 1988).

Reagents for chromosome mapping can be used individually to mark a single chromosome or a single site on that chromosome, or panels of reagents can be used for marking multiple sites and/or multiple chromosomes. Reagents corresponding to noncoding regions of the genes actually are preferred for mapping purposes. Coding

10   sequences are more likely to be conserved within gene families, thus increasing the chance of cross hybridizations during chromosomal mapping.

Once a sequence has been mapped to a precise chromosomal location, the physical position of the sequence on the chromosome can be correlated with genetic map data. Such data are found, *e.g.,* in McKusick, MENDELIAN INHERITANCE IN MAN, available on-line

15   through Johns Hopkins University Welch Medical Library). The relationship between genes and disease, mapped to the same chromosomal region, can then be identified through linkage analysis (co-inheritance of physically adjacent genes), described in, *e.g.,* Egeland, *et al.,* 1987. *Nature,* 325: 783-787.

Moreover, differences in the DNA sequences between individuals affected and

20   unaffected with a disease associated with the NOVX gene, can be determined. If a mutation is observed in some or all of the affected individuals but not in any unaffected individuals, then the mutation is likely to be the causative agent of the particular disease. Comparison of affected and unaffected individuals generally involves first looking for structural alterations in the chromosomes, such as deletions or translocations that are

25   visible from chromosome spreads or detectable using PCR based on that DNA sequence. Ultimately, complete sequencing of genes from several individuals can be performed to confirm the presence of a mutation and to distinguish mutations from polymorphisms.

Tissue Typing

The NOVX sequences of the invention can also be used to identify individuals from

30   minute biological samples. In this technique, an individual's genomic DNA is digested with one or more restriction enzymes, and probed on a Southern blot to yield unique bands for identification. The sequences of the invention are useful as additional DNA markers for

RFLP ("restriction fragment length polymorphisms," described in U.S. Patent No. 5,272,057).

Furthermore, the sequences of the invention can be used to provide an alternative technique that determines the actual base-by-base DNA sequence of selected portions of an

5   individual's genome. Thus, the NOVX sequences described herein can be used to prepare two PCR primers from the 5'- and 3'-termini of the sequences. These primers can then be used to amplify an individual's DNA and subsequently sequence it.

Panels of corresponding DNA sequences from individuals, prepared in this manner, can provide unique individual identifications, as each individual will have a unique set of

10 such DNA sequences due to allelic differences. The sequences of the invention can be used to obtain such identification sequences from individuals and from tissue. The NOVX sequences of the invention uniquely represent portions of the human genome. Allelic variation occurs to some degree in the coding regions of these sequences, and to a greater degree in the noncoding regions. It is estimated that allelic variation between individual

15 humans occurs with a frequency of about once per each 500 bases. Much of the allelic variation is due to single nucleotide polymorphisms (SNPs), which include restriction fragment length polymorphisms (RFLPs).

Each of the sequences described herein can, to some degree, be used as a standard against which DNA from an individual can be compared for identification purposes.

20 Because greater numbers of polymorphisms occur in the noncoding regions, fewer sequences are necessary to differentiate individuals. The noncoding sequences can comfortably provide positive individual identification with a panel of perhaps 10 to 1,000 primers that each yield a noncoding amplified sequence of 100 bases. If coding sequences, such as those of SEQ ID NO:2n-1, wherein n is an integer between 1 and 124, are used, a

25 more appropriate number of primers for positive individual identification would be 500-2,000.

Predictive Medicine

The invention also pertains to the field of predictive medicine in which diagnostic assays, prognostic assays, pharmacogenomics, and monitoring clinical trials are used for

30 prognostic (predictive) purposes to thereby treat an individual prophylactically. Accordingly, one aspect of the invention relates to diagnostic assays for determining NOVX protein and/or nucleic acid expression as well as NOVX activity, in the context of a biological sample (e.g., blood, serum, cells, tissue) to thereby determine whether an

individual is afflicted with a disease or disorder, or is at risk of developing a disorder, associated with aberrant NOVX expression or activity. The disorders include metabolic disorders, diabetes, obesity, infectious disease, anorexia, cancer-associated cachexia, cancer, neurodegenerative disorders, Alzheimer's Disease, Parkinson's Disorder, immune

5   disorders, and hematopoietic disorders, and the various dyslipidemias, metabolic disturbances associated with obesity, the metabolic syndrome X and wasting disorders associated with chronic diseases and various cancers. The invention also provides for prognostic (or predictive) assays for determining whether an individual is at risk of developing a disorder associated with NOVX protein, nucleic acid expression or activity.

10   For example, mutations in a NOVX gene can be assayed in a biological sample. Such assays can be used for prognostic or predictive purpose to thereby prophylactically treat an individual prior to the onset of a disorder characterized by or associated with NOVX protein, nucleic acid expression, or biological activity.

Another aspect of the invention provides methods for determining NOVX protein,

15   nucleic acid expression or activity in an individual to thereby select appropriate therapeutic or prophylactic agents for that individual (referred to herein as "pharmacogenomics"). Pharmacogenomics allows for the selection of agents (*e.g.*, drugs) for therapeutic or prophylactic treatment of an individual based on the genotype of the individual (*e.g.*, the genotype of the individual examined to determine the ability of the individual to respond to

20   a particular agent.)

Yet another aspect of the invention pertains to monitoring the influence of agents (*e.g.*, drugs, compounds) on the expression or activity of NOVX in clinical trials.

These and other agents are described in further detail in the following sections.

Diagnostic Assays

25   An exemplary method for detecting the presence or absence of NOVX in a biological sample involves obtaining a biological sample from a test subject and contacting the biological sample with a compound or an agent capable of detecting NOVX protein or nucleic acid (*e.g.*, mRNA, genomic DNA) that encodes NOVX protein such that the presence of NOVX is detected in the biological sample. An agent for detecting NOVX

30   mRNA or genomic DNA is a labeled nucleic acid probe capable of hybridizing to NOVX mRNA or genomic DNA. The nucleic acid probe can be, for example, a full-length NOVX nucleic acid, such as the nucleic acid of SEQ ID NO:$2n$-1, wherein n is an integer between 1 and 124, or a portion thereof, such as an oligonucleotide of at least 15, 30, 50, 100, 250 or

500 nucleotides in length and sufficient to specifically hybridize under stringent conditions
to NOVX mRNA or genomic DNA. Other suitable probes for use in the diagnostic assays
of the invention are described herein.

An agent for detecting NOVX protein is an antibody capable of binding to NOVX
protein, preferably an antibody with a detectable label. Antibodies can be polyclonal, or
more preferably, monoclonal. An intact antibody, or a fragment thereof (e.g., Fab or
F(ab')₂) can be used. The term "labeled", with regard to the probe or antibody, is intended
to encompass direct labeling of the probe or antibody by coupling (i.e., physically linking)
a detectable substance to the probe or antibody, as well as indirect labeling of the probe or
antibody by reactivity with another reagent that is directly labeled. Examples of indirect
labeling include detection of a primary antibody using a fluorescently-labeled secondary
antibody and end-labeling of a DNA probe with biotin such that it can be detected with
fluorescently-labeled streptavidin. The term "biological sample" is intended to include
tissues, cells and biological fluids isolated from a subject, as well as tissues, cells and fluids
present within a subject. That is, the detection method of the invention can be used to
detect NOVX mRNA, protein, or genomic DNA in a biological sample in vitro as well as
in vivo. For example, in vitro techniques for detection of NOVX mRNA include Northern
hybridizations and in situ hybridizations. In vitro techniques for detection of NOVX
protein include enzyme linked immunosorbent assays (ELISAs), Western blots,
immunoprecipitations, and immunofluorescence. In vitro techniques for detection of
NOVX genomic DNA include Southern hybridizations. Furthermore, in vivo techniques
for detection of NOVX protein include introducing into a subject a labeled anti-NOVX
antibody. For example, the antibody can be labeled with a radioactive marker whose
presence and location in a subject can be detected by standard imaging techniques.

In one embodiment, the biological sample contains protein molecules from the test
subject. Alternatively, the biological sample can contain mRNA molecules from the test
subject or genomic DNA molecules from the test subject. A preferred biological sample is
a peripheral blood leukocyte sample isolated by conventional means from a subject.

In another embodiment, the methods further involve obtaining a control biological
sample from a control subject, contacting the control sample with a compound or agent
capable of detecting NOVX protein, mRNA, or genomic DNA, such that the presence of
NOVX protein, mRNA or genomic DNA is detected in the biological sample, and

comparing the presence of NOVX protein, mRNA or genomic DNA in the control sample with the presence of NOVX protein, mRNA or genomic DNA in the test sample.

The invention also encompasses kits for detecting the presence of NOVX in a biological sample. For example, the kit can comprise: a labeled compound or agent
5   capable of detecting NOVX protein or mRNA in a biological sample; means for determining the amount of NOVX in the sample; and means for comparing the amount of NOVX in the sample with a standard. The compound or agent can be packaged in a suitable container. The kit can further comprise instructions for using the kit to detect NOVX protein or nucleic acid.

10   **Prognostic Assays**

The diagnostic methods described herein can furthermore be utilized to identify subjects having or at risk of developing a disease or disorder associated with aberrant NOVX expression or activity. For example, the assays described herein, such as the preceding diagnostic assays or the following assays, can be utilized to identify a subject
15   having or at risk of developing a disorder associated with NOVX protein, nucleic acid expression or activity. Alternatively, the prognostic assays can be utilized to identify a subject having or at risk for developing a disease or disorder. Thus, the invention provides a method for identifying a disease or disorder associated with aberrant NOVX expression or activity in which a test sample is obtained from a subject and NOVX protein or nucleic
20   acid (*e.g.*, mRNA, genomic DNA) is detected, wherein the presence of NOVX protein or nucleic acid is diagnostic for a subject having or at risk of developing a disease or disorder associated with aberrant NOVX expression or activity. As used herein, a "test sample" refers to a biological sample obtained from a subject of interest. For example, a test sample can be a biological fluid (*e.g.*, serum), cell sample, or tissue.

25   Furthermore, the prognostic assays described herein can be used to determine whether a subject can be administered an agent (*e.g.*, an agonist, antagonist, peptidomimetic, protein, peptide, nucleic acid, small molecule, or other drug candidate) to treat a disease or disorder associated with aberrant NOVX expression or activity. For example, such methods can be used to determine whether a subject can be effectively
30   treated with an agent for a disorder. Thus, the invention provides methods for determining whether a subject can be effectively treated with an agent for a disorder associated with aberrant NOVX expression or activity in which a test sample is obtained and NOVX protein or nucleic acid is detected (*e.g.*, wherein the presence of NOVX protein or nucleic

acid is diagnostic for a subject that can be administered the agent to treat a disorder associated with aberrant NOVX expression or activity).

The methods of the invention can also be used to detect genetic lesions in a NOVX gene, thereby determining if a subject with the lesioned gene is at risk for a disorder

5   characterized by aberrant cell proliferation and/or differentiation. In various embodiments, the methods include detecting, in a sample of cells from the subject, the presence or absence of a genetic lesion characterized by at least one of an alteration affecting the integrity of a gene encoding a NOVX-protein, or the misexpression of the NOVX gene. For example, such genetic lesions can be detected by ascertaining the existence of at least

10   one of: (*i*) a deletion of one or more nucleotides from a NOVX gene; (*ii*) an addition of one or more nucleotides to a NOVX gene; (*iii*) a substitution of one or more nucleotides of a NOVX gene, (*iv*) a chromosomal rearrangement of a NOVX gene; (*v*) an alteration in the level of a messenger RNA transcript of a NOVX gene, (*vi*) aberrant modification of a NOVX gene, such as of the methylation pattern of the genomic DNA, (*vii*) the presence of

15   a non-wild-type splicing pattern of a messenger RNA transcript of a NOVX gene, (*viii*) a non-wild-type level of a NOVX protein, (*ix*) allelic loss of a NOVX gene, and (*x*) inappropriate post-translational modification of a NOVX protein. As described herein, there are a large number of assay techniques known in the art which can be used for detecting lesions in a NOVX gene. A preferred biological sample is a peripheral blood

20   leukocyte sample isolated by conventional means from a subject. However, any biological sample containing nucleated cells may be used, including, for example, buccal mucosal cells.

In certain embodiments, detection of the lesion involves the use of a probe/primer in a polymerase chain reaction (PCR) (*see, e.g.*, U.S. Patent Nos. 4,683,195 and 4,683,202),

25   such as anchor PCR or RACE PCR, or, alternatively, in a ligation chain reaction (LCR) (*see, e.g.*, Landegran, *et al.*, 1988. *Science* 241: 1077-1080; and Nakazawa, *et al.*, 1994. *Proc. Natl. Acad. Sci. USA* 91: 360-364), the latter of which can be particularly useful for detecting point mutations in the NOVX-gene (*see*, Abravaya, *et al.*, 1995. *Nucl. Acids Res.* 23: 675-682). This method can include the steps of collecting a sample of cells from a

30   patient, isolating nucleic acid (*e.g.*, genomic, mRNA or both) from the cells of the sample, contacting the nucleic acid sample with one or more primers that specifically hybridize to a NOVX gene under conditions such that hybridization and amplification of the NOVX gene (if present) occurs, and detecting the presence or absence of an amplification product, or

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detecting the size of the amplification product and comparing the length to a control sample. It is anticipated that PCR and/or LCR may be desirable to use as a preliminary amplification step in conjunction with any of the techniques used for detecting mutations described herein.

5   Alternative amplification methods include: self sustained sequence replication (*see*, Guatelli, *et al.*, 1990. *Proc. Natl. Acad. Sci. USA* 87: 1874-1878), transcriptional amplification system (*see*, Kwoh, *et al.*, 1989. *Proc. Natl. Acad. Sci. USA* 86: 1173-1177); Qβ Replicase (*see*, Lizardi, *et al*, 1988. *BioTechnology* 6: 1197), or any other nucleic acid amplification method, followed by the detection of the amplified molecules using

10   techniques well known to those of skill in the art. These detection schemes are especially useful for the detection of nucleic acid molecules if such molecules are present in very low numbers.

In an alternative embodiment, mutations in a NOVX gene from a sample cell can be identified by alterations in restriction enzyme cleavage patterns. For example, sample and

15   control DNA is isolated, amplified (optionally), digested with one or more restriction endonucleases, and fragment length sizes are determined by gel electrophoresis and compared. Differences in fragment length sizes between sample and control DNA indicates mutations in the sample DNA. Moreover, the use of sequence specific ribozymes (*see, e.g.,* U.S. Patent No. 5,493,531) can be used to score for the presence of specific

20   mutations by development or loss of a ribozyme cleavage site.

In other embodiments, genetic mutations in NOVX can be identified by hybridizing a sample and control nucleic acids, *e.g.*, DNA or RNA, to high-density arrays containing hundreds or thousands of oligonucleotides probes. *See, e.g.*, Cronin, *et al.*, 1996. *Human Mutation* 7: 244-255; Kozal, *et al.*, 1996. *Nat. Med.* 2: 753-759. For example, genetic

25   mutations in NOVX can be identified in two dimensional arrays containing light-generated DNA probes as described in Cronin, *et al.*, *supra*. Briefly, a first hybridization array of probes can be used to scan through long stretches of DNA in a sample and control to identify base changes between the sequences by making linear arrays of sequential overlapping probes. This step allows the identification of point mutations. This is

30   followed by a second hybridization array that allows the characterization of specific mutations by using smaller, specialized probe arrays complementary to all variants or mutations detected. Each mutation array is composed of parallel probe sets, one complementary to the wild-type gene and the other complementary to the mutant gene.

In yet another embodiment, any of a variety of sequencing reactions known in the art can be used to directly sequence the NOVX gene and detect mutations by comparing the sequence of the sample NOVX with the corresponding wild-type (control) sequence. Examples of sequencing reactions include those based on techniques developed by Maxim and Gilbert, 1977. *Proc. Natl. Acad. Sci. USA* 74: 560 or Sanger, 1977. *Proc. Natl. Acad. Sci. USA* 74: 5463. It is also contemplated that any of a variety of automated sequencing procedures can be utilized when performing the diagnostic assays (*see, e.g.,* Naeve, *et al.,* 1995. *Biotechniques* 19: 448), including sequencing by mass spectrometry (see, *e.g.,* PCT International Publication No. WO 94/16101; Cohen, *et al.,* 1996. *Adv. Chromatography* 36: 127-162; and Griffin, *et al.,* 1993. *Appl. Biochem. Biotechnol.* 38: 147-159).

Other methods for detecting mutations in the NOVX gene include methods in which protection from cleavage agents is used to detect mismatched bases in RNA/RNA or RNA/DNA heteroduplexes. *See, e.g.,* Myers, *et al.,* 1985. *Science* 230: 1242. In general, the art technique of "mismatch cleavage" starts by providing heteroduplexes of formed by hybridizing (labeled) RNA or DNA containing the wild-type NOVX sequence with potentially mutant RNA or DNA obtained from a tissue sample. The double-stranded duplexes are treated with an agent that cleaves single-stranded regions of the duplex such as which will exist due to basepair mismatches between the control and sample strands. For instance, RNA/DNA duplexes can be treated with RNase and DNA/DNA hybrids treated with S_1 nuclease to enzymatically digesting the mismatched regions. In other embodiments, either DNA/DNA or RNA/DNA duplexes can be treated with hydroxylamine or osmium tetroxide and with piperidine in order to digest mismatched regions. After digestion of the mismatched regions, the resulting material is then separated by size on denaturing polyacrylamide gels to determine the site of mutation. *See, e.g.,* Cotton, *et al.,* 1988. *Proc. Natl. Acad. Sci. USA* 85: 4397; Saleeba, *et al.,* 1992. *Methods Enzymol.* 217: 286-295. In an embodiment, the control DNA or RNA can be labeled for detection.

In still another embodiment, the mismatch cleavage reaction employs one or more proteins that recognize mismatched base pairs in double-stranded DNA (so called "DNA mismatch repair" enzymes) in defined systems for detecting and mapping point mutations in NOVX cDNAs obtained from samples of cells. For example, the mutY enzyme of *E. coli* cleaves A at G/A mismatches and the thymidine DNA glycosylase from HeLa cells cleaves T at G/T mismatches. *See, e.g.,* Hsu, *et al.,* 1994. *Carcinogenesis* 15: 1657-1662.

According to an exemplary embodiment, a probe based on a NOVX sequence, e.g., a wild-type NOVX sequence, is hybridized to a cDNA or other DNA product from a test cell(s). The duplex is treated with a DNA mismatch repair enzyme, and the cleavage products, if any, can be detected from electrophoresis protocols or the like. See, e.g., U.S.

5   Patent No. 5,459,039.

In other embodiments, alterations in electrophoretic mobility will be used to identify mutations in NOVX genes. For example, single strand conformation polymorphism (SSCP) may be used to detect differences in electrophoretic mobility between mutant and wild type nucleic acids. See, e.g., Orita, et al., 1989. Proc. Natl. Acad.

10   Sci. USA: 86: 2766; Cotton, 1993. Mutat. Res. 285: 125-144; Hayashi, 1992. Genet. Anal. Tech. Appl. 9: 73-79. Single-stranded DNA fragments of sample and control NOVX nucleic acids will be denatured and allowed to renature. The secondary structure of single-stranded nucleic acids varies according to sequence, the resulting alteration in electrophoretic mobility enables the detection of even a single base change. The DNA

15   fragments may be labeled or detected with labeled probes. The sensitivity of the assay may be enhanced by using RNA (rather than DNA), in which the secondary structure is more sensitive to a change in sequence. In one embodiment, the subject method utilizes heteroduplex analysis to separate double stranded heteroduplex molecules on the basis of changes in electrophoretic mobility. See, e.g., Keen, et al., 1991. Trends Genet. 7: 5.

20   In yet another embodiment, the movement of mutant or wild-type fragments in polyacrylamide gels containing a gradient of denaturant is assayed using denaturing gradient gel electrophoresis (DGGE). See, e.g., Myers, et al., 1985. Nature 313: 495. When DGGE is used as the method of analysis, DNA will be modified to insure that it does not completely denature, for example by adding a GC clamp of approximately 40 bp of

25   high-melting GC-rich DNA by PCR. In a further embodiment, a temperature gradient is used in place of a denaturing gradient to identify differences in the mobility of control and sample DNA. See, e.g., Rosenbaum and Reissner, 1987. Biophys. Chem. 265: 12753.

Examples of other techniques for detecting point mutations include, but are not limited to, selective oligonucleotide hybridization, selective amplification, or selective

30   primer extension. For example, oligonucleotide primers may be prepared in which the known mutation is placed centrally and then hybridized to target DNA under conditions that permit hybridization only if a perfect match is found. See, e.g., Saiki, et al., 1986. Nature 324: 163; Saiki, et al., 1989. Proc. Natl. Acad. Sci. USA 86: 6230. Such allele

specific oligonucleotides are hybridized to PCR amplified target DNA of a number of different mutations when the oligonucleotides are attached to the hybridizing membrane and hybridized with labeled target DNA.

Alternatively, allele specific amplification technology that depends on selective PCR amplification may be used in conjunction with the instant invention. Oligonucleotides used as primers for specific amplification may carry the mutation of interest in the center of the molecule (so that amplification depends on differential hybridization; *see, e.g.,* Gibbs, *et al.,* 1989. *Nucl. Acids Res.* 17: 2437-2448) or at the extreme 3'-terminus of one primer where, under appropriate conditions, mismatch can prevent, or reduce polymerase extension (*see, e.g.,* Prossner, 1993. *Tibtech.* 11: 238). In addition it may be desirable to introduce a novel restriction site in the region of the mutation to create cleavage-based detection. *See, e.g.,* Gasparini, *et al.,* 1992. *Mol. Cell Probes* 6: 1. It is anticipated that in certain embodiments amplification may also be performed using *Taq* ligase for amplification. *See, e.g.,* Barany, 1991. *Proc. Natl. Acad. Sci. USA* 88: 189. In such cases, ligation will occur only if there is a perfect match at the 3'-terminus of the 5' sequence, making it possible to detect the presence of a known mutation at a specific site by looking for the presence or absence of amplification.

The methods described herein may be performed, for example, by utilizing pre-packaged diagnostic kits comprising at least one probe nucleic acid or antibody reagent described herein, which may be conveniently used, *e.g.,* in clinical settings to diagnose patients exhibiting symptoms or family history of a disease or illness involving a NOVX gene.

Furthermore, any cell type or tissue, preferably peripheral blood leukocytes, in which NOVX is expressed may be utilized in the prognostic assays described herein. However, any biological sample containing nucleated cells may be used, including, for example, buccal mucosal cells.

Pharmacogenomics

Agents, or modulators that have a stimulatory or inhibitory effect on NOVX activity (*e.g.,* NOVX gene expression), as identified by a screening assay described herein can be administered to individuals to treat (prophylactically or therapeutically) disorders. The disorders include but are not limited to, *e.g.,* those diseases, disorders and conditions listed above, and more particularly include those diseases, disorders, or conditions associated with homologs of a NOVX protein, such as those summarized in Table A.

In conjunction with such treatment, the pharmacogenomics (i.e., the study of the relationship between an individual's genotype and that individual's response to a foreign compound or drug) of the individual may be considered. Differences in metabolism of therapeutics can lead to severe toxicity or therapeutic failure by altering the relation

5   between dose and blood concentration of the pharmacologically active drug. Thus, the pharmacogenomics of the individual permits the selection of effective agents (e.g., drugs) for prophylactic or therapeutic treatments based on a consideration of the individual's genotype. Such pharmacogenomics can further be used to determine appropriate dosages and therapeutic regimens. Accordingly, the activity of NOVX protein, expression of

10   NOVX nucleic acid, or mutation content of NOVX genes in an individual can be determined to thereby select appropriate agent(s) for therapeutic or prophylactic treatment of the individual.

        Pharmacogenomics deals with clinically significant hereditary variations in the response to drugs due to altered drug disposition and abnormal action in affected persons.

15   See e.g., Eichelbaum, 1996. Clin. Exp. Pharmacol. Physiol., 23: 983-985; Linder, 1997. Clin. Chem., 43: 254-266. In general, two types of pharmacogenetic conditions can be differentiated. Genetic conditions transmitted as a single factor altering the way drugs act on the body (altered drug action) or genetic conditions transmitted as single factors altering the way the body acts on drugs (altered drug metabolism). These pharmacogenetic

20   conditions can occur either as rare defects or as polymorphisms. For example, glucose-6-phosphate dehydrogenase (G6PD) deficiency is a common inherited enzymopathy in which the main clinical complication is hemolysis after ingestion of oxidant drugs (anti-malarials, sulfonamides, analgesics, nitrofurans) and consumption of fava beans.

25   As an illustrative embodiment, the activity of drug metabolizing enzymes is a major determinant of both the intensity and duration of drug action. The discovery of genetic polymorphisms of drug metabolizing enzymes (e.g., N-acetyltransferase 2 (NAT 2) and cytochrome pregnancy zone protein precursor enzymes CYP2D6 and CYP2C19) has provided an explanation as to why some patients do not obtain the expected drug effects or

30   show exaggerated drug response and serious toxicity after taking the standard and safe dose of a drug. These polymorphisms are expressed in two phenotypes in the population, the extensive metabolizer (EM) and poor metabolizer (PM). The prevalence of PM is different among different populations. For example, the gene coding for CYP2D6 is highly

polymorphic and several mutations have been identified in PM, which all lead to the absence of functional CYP2D6. Poor metabolizers of CYP2D6 and CYP2C19 quite frequently experience exaggerated drug response and side effects when they receive standard doses. If a metabolite is the active therapeutic moiety, PM show no therapeutic

5   response, as demonstrated for the analgesic effect of codeine mediated by its CYP2D6-formed metabolite morphine. At the other extreme are the so called ultra-rapid metabolizers who do not respond to standard doses. Recently, the molecular basis of ultra-rapid metabolism has been identified to be due to CYP2D6 gene amplification.

Thus, the activity of NOVX protein, expression of NOVX nucleic acid, or mutation

10   content of NOVX genes in an individual can be determined to thereby select appropriate agent(s) for therapeutic or prophylactic treatment of the individual. In addition, pharmacogenetic studies can be used to apply genotyping of polymorphic alleles encoding drug-metabolizing enzymes to the identification of an individual's drug responsiveness phenotype. This knowledge, when applied to dosing or drug selection, can avoid adverse

15   reactions or therapeutic failure and thus enhance therapeutic or prophylactic efficiency when treating a subject with a NOVX modulator, such as a modulator identified by one of the exemplary screening assays described herein.

Monitoring of Effects During Clinical Trials

Monitoring the influence of agents (e.g., drugs, compounds) on the expression or

20   activity of NOVX (e.g., the ability to modulate aberrant cell proliferation and/or differentiation) can be applied not only in basic drug screening, but also in clinical trials. For example, the effectiveness of an agent determined by a screening assay as described herein to increase NOVX gene expression, protein levels, or upregulate NOVX activity, can be monitored in clinical trails of subjects exhibiting decreased NOVX gene expression,

25   protein levels, or downregulated NOVX activity. Alternatively, the effectiveness of an agent determined by a screening assay to decrease NOVX gene expression, protein levels, or downregulate NOVX activity, can be monitored in clinical trails of subjects exhibiting increased NOVX gene expression, protein levels, or upregulated NOVX activity. In such clinical trials, the expression or activity of NOVX and, preferably, other genes that have

30   been implicated in, for example, a cellular proliferation or immune disorder can be used as a "read out" or markers of the immune responsiveness of a particular cell.

By way of example, and not of limitation, genes, including NOVX, that are modulated in cells by treatment with an agent (e.g., compound, drug or small molecule)

that modulates NOVX activity (*e.g.*, identified in a screening assay as described herein) can be identified. Thus, to study the effect of agents on cellular proliferation disorders, for example, in a clinical trial, cells can be isolated and RNA prepared and analyzed for the levels of expression of NOVX and other genes implicated in the disorder. The levels of

5   gene expression (*i.e.*, a gene expression pattern) can be quantified by Northern blot analysis or RT-PCR, as described herein, or alternatively by measuring the amount of protein produced, by one of the methods as described herein, or by measuring the levels of activity of NOVX or other genes. In this manner, the gene expression pattern can serve as a marker, indicative of the physiological response of the cells to the agent. Accordingly, this

10   response state may be determined before, and at various points during, treatment of the individual with the agent.

In one embodiment, the invention provides a method for monitoring the effectiveness of treatment of a subject with an agent (*e.g.*, an agonist, antagonist, protein, peptide, peptidomimetic, nucleic acid, small molecule, or other drug candidate identified by

15   the screening assays described herein) comprising the steps of (*i*) obtaining a pre-administration sample from a subject prior to administration of the agent; (*ii*) detecting the level of expression of a NOVX protein, mRNA, or genomic DNA in the preadministration sample; (*iii*) obtaining one or more post-administration samples from the subject; (*iv*) detecting the level of expression or activity of the NOVX protein, mRNA, or

20   genomic DNA in the post-administration samples; (*v*) comparing the level of expression or activity of the NOVX protein, mRNA, or genomic DNA in the pre-administration sample with the NOVX protein, mRNA, or genomic DNA in the post administration sample or samples; and (*vi*) altering the administration of the agent to the subject accordingly. For example, increased administration of the agent may be desirable to increase the expression

25   or activity of NOVX to higher levels than detected, *i.e.*, to increase the effectiveness of the agent. Alternatively, decreased administration of the agent may be desirable to decrease expression or activity of NOVX to lower levels than detected, *i.e.*, to decrease the effectiveness of the agent.

Methods of Treatment

30   The invention provides for both prophylactic and therapeutic methods of treating a subject at risk of (or susceptible to) a disorder or having a disorder associated with aberrant NOVX expression or activity. The disorders include but are not limited to, *e.g.*, those diseases, disorders and conditions listed above, and more particularly include those

diseases, disorders, or conditions associated with homologs of a NOVX protein, such as those summarized in Table A.

These methods of treatment will be discussed more fully, below.

Diseases and Disorders

5   Diseases and disorders that are characterized by increased (relative to a subject not suffering from the disease or disorder) levels or biological activity may be treated with Therapeutics that antagonize (*i.e.*, reduce or inhibit) activity. Therapeutics that antagonize activity may be administered in a therapeutic or prophylactic manner. Therapeutics that may be utilized include, but are not limited to: (*i*) an aforementioned peptide, or analogs,

10   derivatives, fragments or homologs thereof; (*ii*) antibodies to an aforementioned peptide; (*iii*) nucleic acids encoding an aforementioned peptide; (*iv*) administration of antisense nucleic acid and nucleic acids that are "dysfunctional" (*i.e.*, due to a heterologous insertion within the coding sequences of coding sequences to an aforementioned peptide) that are utilized to "knockout" endogenous function of an aforementioned peptide by homologous

15   recombination (*see, e.g.,* Capecchi, 1989. *Science* 244: 1288-1292); or (*v*) modulators (*i.e.*, inhibitors, agonists and antagonists, including additional peptide mimetic of the invention or antibodies specific to a peptide of the invention) that alter the interaction between an aforementioned peptide and its binding partner.

Diseases and disorders that are characterized by decreased (relative to a subject not

20   suffering from the disease or disorder) levels or biological activity may be treated with Therapeutics that increase (*i.e.*, are agonists to) activity. Therapeutics that upregulate activity may be administered in a therapeutic or prophylactic manner. Therapeutics that may be utilized include, but are not limited to, an aforementioned peptide, or analogs, derivatives, fragments or homologs thereof; or an agonist that increases bioavailability.

25   Increased or decreased levels can be readily detected by quantifying peptide and/or RNA, by obtaining a patient tissue sample (*e.g.,* from biopsy tissue) and assaying it *in vitro* for RNA or peptide levels, structure and/or activity of the expressed peptides (or mRNAs of an aforementioned peptide). Methods that are well-known within the art include, but are not limited to, immunoassays (*e.g.,* by Western blot analysis, immunoprecipitation

30   followed by sodium dodecyl sulfate (SDS) polyacrylamide gel electrophoresis, immunocytochemistry, *etc.*) and/or hybridization assays to detect expression of mRNAs (*e.g.,* Northern assays, dot blots, *in situ* hybridization, and the like).

Prophylactic Methods

In one aspect, the invention provides a method for preventing, in a subject, a disease or condition associated with an aberrant NOVX expression or activity, by administering to the subject an agent that modulates NOVX expression or at least one NOVX activity.

5 Subjects at risk for a disease that is caused or contributed to by aberrant NOVX expression or activity can be identified by, for example, any or a combination of diagnostic or prognostic assays as described herein. Administration of a prophylactic agent can occur prior to the manifestation of symptoms characteristic of the NOVX aberrancy, such that a disease or disorder is prevented or, alternatively, delayed in its progression. Depending

10 upon the type of NOVX aberrancy, for example, a NOVX agonist or NOVX antagonist agent can be used for treating the subject. The appropriate agent can be determined based on screening assays described herein. The prophylactic methods of the invention are further discussed in the following subsections.

Therapeutic Methods

15 Another aspect of the invention pertains to methods of modulating NOVX expression or activity for therapeutic purposes. The modulatory method of the invention involves contacting a cell with an agent that modulates one or more of the activities of NOVX protein activity associated with the cell. An agent that modulates NOVX protein activity can be an agent as described herein, such as a nucleic acid or a protein, a

20 naturally-occurring cognate ligand of a NOVX protein, a peptide, a NOVX peptidomimetic, or other small molecule. In one embodiment, the agent stimulates one or more NOVX protein activity. Examples of such stimulatory agents include active NOVX protein and a nucleic acid molecule encoding NOVX that has been introduced into the cell. In another embodiment, the agent inhibits one or more NOVX protein activity. Examples

25 of such inhibitory agents include antisense NOVX nucleic acid molecules and anti-NOVX antibodies. These modulatory methods can be performed *in vitro* (*e.g.*, by culturing the cell with the agent) or, alternatively, *in vivo* (*e.g.*, by administering the agent to a subject). As such, the invention provides methods of treating an individual afflicted with a disease or disorder characterized by aberrant expression or activity of a NOVX protein or nucleic acid

30 molecule. In one embodiment, the method involves administering an agent (*e.g.*, an agent identified by a screening assay described herein), or combination of agents that modulates (*e.g.*, up-regulates or down-regulates) NOVX expression or activity. In another

embodiment, the method involves administering a NOVX protein or nucleic acid molecule as therapy to compensate for reduced or aberrant NOVX expression or activity.

Stimulation of NOVX activity is desirable *in situ*ations in which NOVX is abnormally downregulated and/or in which increased NOVX activity is likely to have a

5   beneficial effect. One example of such a situation is where a subject has a disorder characterized by aberrant cell proliferation and/or differentiation (*e.g.*, cancer or immune associated disorders). Another example of such a situation is where the subject has a gestational disease (*e.g.*, preclampsia).

Determination of the Biological Effect of the Therapeutic

10   In various embodiments of the invention, suitable *in vitro* or *in vivo* assays are performed to determine the effect of a specific Therapeutic and whether its administration is indicated for treatment of the affected tissue.

In various specific embodiments, *in vitro* assays may be performed with representative cells of the type(s) involved in the patient's disorder, to determine if a given

15   Therapeutic exerts the desired effect upon the cell type(s). Compounds for use in therapy may be tested in suitable animal model systems including, but not limited to rats, mice, chicken, cows, monkeys, rabbits, and the like, prior to testing in human subjects. Similarly, for *in vivo* testing, any of the animal model system known in the art may be used prior to administration to human subjects.

20   **Prophylactic and Therapeutic Uses of the Compositions of the Invention**

The NOVX nucleic acids and proteins of the invention are useful in potential prophylactic and therapeutic applications implicated in a variety of disorders. The disorders include but are not limited to, *e.g.*, those diseases, disorders and conditions listed above, and more particularly include those diseases, disorders, or conditions associated

25   with homologs of a NOVX protein, such as those summarized in Table A.

As an example, a cDNA encoding the NOVX protein of the invention may be useful in gene therapy, and the protein may be useful when administered to a subject in need thereof. By way of non-limiting example, the compositions of the invention will have efficacy for treatment of patients suffering from diseases, disorders, conditions and the like,

30   including but not limited to those listed herein.

Both the novel nucleic acid encoding the NOVX protein, and the NOVX protein of the invention, or fragments thereof, may also be useful in diagnostic applications, wherein

the presence or amount of the nucleic acid or the protein are to be assessed. A further use could be as an anti-bacterial molecule (*i.e.,* some peptides have been found to possess anti-bacterial properties). These materials are further useful in the generation of antibodies, which immunospecifically-bind to the novel substances of the invention for use in

5   therapeutic or diagnostic methods.

The invention will be further described in the following examples, which do not limit the scope of the invention described in the claims.

EXAMPLES

Example A: Polynucleotide and Polypeptide Sequences, and Homology Data

10   **Example 1.**

The NOV1 clone was analyzed, and the nucleotide and encoded polypeptide sequences are shown in Table 1A.

Table 1A. NOV1 Sequence Analysis			
	SEQ ID NO: 1	6189 bp	
NOV1a, CG106764-01 DNA Sequence	ATGTTGAAGTTCAAATATGGAGCGCGGAATCCTTTGGATGCTGGTGCTGCTGAACCCATTGCCAGCCG GGCCTCCAGGCTGAATCTGTTCTTCCAGGGGAAACCACCCTTTATGACTCAACAGCAGATGTCTCCTC TTTCCCGAGAAGGGATATTAGATGCCCTCTTTGTTCTCTTTGAAGAATGCAGTCAGCCTGCTCTGATG AAGATTAAGCACGTGAGCAACTTTGTCCGGAAGTGTTCCGACACCATAGCTGAGTTACAGGAGCTCCA GCCTTCGGCAAAGGACTTCGAAGTCAGAAGTCTTGTAGGTTGTGGTCACTTTGCTGAAGTGCAGGTGG TAAGAGAGAAAGCAACCGGGGACATCTATGCTATGAAAGTGATGAAGAAGAAGGCTTTATTGGCCCAG GAGCAGGTTTCATTTTTTGAGGAAGAGCGGAACATATTATCTCGAAGCACAAGCCCGTGGATCCCCCA ATTACAGTATGCCTTTCAGGACAAAAATCACCTTTATCTGGTGATGGAATATCAGCCTGGAGGGGACT TGCTGTCACTTTTGAATAGATATGAGGACCAGTTAGATGAAAACCTGATACAGTTTTACCTAGCTGAG CTGATTTTGGCTGTTCACAGCGTTCATCTGATGGGATACGTGCATCGGGACATCAAGCCTGAGAACAT TCTCGTTGACCGCACAGGACACATCAAGCTGGTGGATTTTGGATCTGCCGCGAAAATGAATTCAAACA AGGTGAATGCCAAACTCCCGATTGGGACCCCAGATTACATGGCTCCTGAAGTGCTGACTGTGATGAAC GGGGATGGGAAAAGGCACCTACGGCCTGGACTGTGCACTGGTGGTCAGTGGGCGTGATTGCCTATGAGAT GATTTATGGGAGATCCCCCTTCGCAGAGGGAACCTCTGCCAGAACCTTCAATAACATTATGAATTTCC AGCGGTTTTTGAAATTTCCAGATGACCCCAAAGTGAGCAGTGACTTTCTTGATCTGATTCAAAGCTTG TTGTGCGGCCAGAAAGAGAGACTGAAGTTTGAAGGTCTTTGCTGCCATCCTTTCTTTCTCTAAAATTGA CTGGAACAACATTCGTAACGCTCCTCCCCCCTTCGTTCCCACCCTCAAGTCTGACGATGACACCTCCA ATTTTGATGAACCAGAGAAGAATTCGTGGGTTTCATCCTCTCCGTGCCAGCTGAGCCCCTCAGGCTTC TCGGGTGAAGAACTGCCGTTTGTGGGGTTTTCGTACAGCCAAGGCACTGGGGATTCTTGGTAGATCTGA GTCTGTTGTGTCGGGTCTGGACTCCCCTGCCAAGACTAGCTCCATGGAAAAGAAACTTCTCATCAAAA GCAAAGAGCTACAAGACTCTCAGGACAAGTGTCACAAGATGGAGCAGGAAATGACCCGGTTACATCGG AGAGTGTCAGAGGTGGGAGGCTGTGCTTAGTCAGAAGGAGGTGGAGCTGAAGGCCTCTGAGACTCAGAG ATCCCTCCTGGAGCAGGACCTTGCTACCTACATCACAGAATGCAGTAGCTTAAAGCGAAGTTTGGAGC AAGCACGGATGGAGGTGTCCCAGGAGGATGACAAAGCACTGCAGCTTCTCCATGATATCAGAGAGCAG AGCCGGAAGCTCCAAGAAATCAAAGAGCAGGAGTACCAGGCTCAAGTGGAAGAAATGAGGTTGATGAT GAATCAGTTGGAAGAGGATCTTGTCTCAGCAAGGAACGGGAGTGATCTCTACGAATCTGAGCTGAGAG AGTCTCGGCTTGCTGCTGAAGAATTCAAGCGGAAAGCGACAGAATGTCAGCATAAACTGTTGAAGGCT AAGGATCAGGGGAAGCCTGAAGTGGGGAGAATATGCGAAACTGGAGAAGATCAATGCTGAGCAGCAGCT CAAAATTCAGGAGCTCCAAGAGAAACTGGAGAAGGCTGTAAAAGCCAGCACGGAGGCCACCGAGCTGC TGCAGAATATCCGCCAGGCAAAGGAGCGAGCCGAGAGGGAGCTGGAGAAGCTGCAGAACCGAGAGGAT TCTTCTGAAGGCATCAGAAAGAAGCTGGTGGAAGCTGAGGAACGCCGCCATTCTCTGGAGAACAAGGT AAAGAGACTAGAGACCATGGAGCGTAGAGAAAACAGACTGAAGGATGACATCCAGACAAAATCCCAAC AGATCCAGCAGATGGCTGATAAAATTCTGGAGCTCGAAGAGAAACATCGGGAGGCCCAAGTCTCAGCC CAGCACCTAGAAGTGCACCTGAAACAGAAAGAGCAGCACTATGAGGAAAAGATTAAAGTATTGGACAA TCAGATAAAGAAAGACCTGGCTGACAAGGAGACACTGGAGAACATGATGCAGAGACACGAGGAGGAGG CCCATGAGAAGGGCAAAATTCTCAGCGAACAGAAGGCGATGATCAATGCTATGGATTCCAAGATCAGA TCCCTGGAACAGAGGATTGTGGAACTGTCTGAAGCCAATAAACTTGCAGCAAATAGCAGTCTTTTTAC CCAAAGGAACATGAAGGCCCAAGAAGAGATGATTTCTGAACTCAGGCAACAGAAATTTTACCTGGAGA

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CACAGGCTGGGAAGTTGGAGGCCCAGAACCGAAAACTGGAGGAGCAGCTGGAGAAGATCAGCCACCAA
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GGAGCAGAAACTGGAGCTCAAGCGCCAGCTCACAGAGCTACAGCTCTCCCTGCAGGAGCGCGAGTCAC
AGTTGACAGCCCTGCAGGCTGCACGGGCGGCCCTGGAGAGCCAGCTTCGCCAGGCGAAGACAGAGCTG
GAAGAGACCACAGCAGAAGCTGAAGAGGAGATCCAGGCACTCACGGCACATAGAGATGAAATCCAGCG
CAAATTTGATGCTCTTCGTAACAGCTGTACTGTGATCACAGACCTGGAGGAGCAGCTAAACCAGCTGA
CCGAGGACAACGCTGAACTCAACAACCAAACTTCTACTTGTCCAAACAACTCGATGAGGCTTCTGGC
GCCAACGACGAGATTGTACAACTGCGAAGTGAAGTGGACCATCTCCGCCGGGAGATCACGGAACGAGA
GATGCAGCTTACCAGCCAGAAGCAAACGATGGAGGCTCTGAAGACCACGTGCACCATGCTGGAGGAAC
AGGTCATGGATTTGGAGGCCCTAAACGATGAGCTGCTAGAAAAAGAGCGGCAGTGGGAGGCCTGGAGG
AGCGTCCTGGGTGATGAGAAATCCCAGTTTGAGTGTCGGGTTCGAGAGCTGCAGAGGATGCTGGACAC
CGAGAAACAGAGCAGGGCGAGAGCCGATCAGCGGATCACCGAGTCTCGCCAGGTGGTGGAGCTGGCAG
TGAAGGAGCACAAGGCTGAGATTCTCGCTCTGCAGCAGGCTCTCAAAGAGCAGAAGCTGAAGGCCGAG
AGCCTCTCTGACAAGCTCAATGACCTGGAGAAGAAGCATGCTATGCTTGAAATGAATGCCCGAAGCTT
ACAGCAGAAGCTGGAGACTGAACGAGAGCTCAAACAGAGGCTTCTGGAAGAGCAAGCCAAATTACAGC
AGCAGATGGACCTGCAGAAAAATCACATTTTCCGTCTGACTCAAGGACTGCAAGAAGCTCTAGATCGG
GCTGATCTACTGAAGACAGAAAGAAGTGACTTGGAGTATCAGCTGGAAAACATTCAGGTGCTCTATTC
TCATGAAAAGGTGAAAATGGAAGGCAACTATTTCTCAACAAACCAAACTCATTGATTTTCTGCAAGCCA
AAATGGACCAACCTGCTAAAAAGAAAAAGGTGCCTCTGCAGTACAATGAGCTGAAGCTGGCCCTGGAG
AAGGAGAAAGCTCGCTGTGCAGAGCTAGAGGAAGCCCTTCAGAAGACCCGCATCGAGCTCCGGTCCGC
CCGGGAGGAAGCTGCCCACCGCAAAGCAACGGACCACCCACACCCATCCACGCCAGCCACCGCGAGGC
AGCAGATCGCCATGTCTGCCATCGTGCGGTCGTGCGGTCGCCAGAGCACCAGCCCAGTGCCATGAGCCTGCTGGCC
CCGCCATCCAGCCGCAGAAAGGAGTCTTCAACTCCAGAGGAATTTAGTCGGCGTCTTAAGGAACGCAT
GCACCACAATATTCCTCACCGATTCAACGTAGGACTGAACATGCGAGCCACAAAGTGTGCTGTGTGTC
TGGATACCGTGCACTTTGGACGCCAGGCATCCAAATGTCTAGAATGTCAGGTGATGTGTCACCCCAAG
TGCTCCACGTGCTTGCCAGCCACCTGCGGCTTGCCTGCTGAATATGCCACACACTTCACCGAGGCCTT
CTGCCGTGACAAAATGAACTCCCCAGGTCTCCAGACCAAGGAGCCCAGCAGCAGCTTGCACCTGGAAG
GGTGGATGAAGGTGCCCAGGAATAACAAACGAGGACAGCAAGGCTGGGACAGGAAGTACATTGTCCTG
GAGGGATCAAAAGTCCTCATTTATGACAATGAAGCCAGAGAAGCTGGACAGAGGCCGGTGGAAGAATT
TGAGCTGTGCCTTCCCGACGGGGATGTATCTATTCATGGTGCCGTTGGTGCTTCCGAACTCGCAAATA
CAGCCAAAGCAGATGTCCCATACATACTGAAGATGGAATCTCACCCGCACACCACCTGCTGGCCCGGG
AGAACCCTCTACTTGCTAGCTCCCAGCTTCCCTGACAAACAGCGCTGGGTCACCGCCTTAGAATCAGT
TGTCGCAGGTGGGAGAGTTTCTAGGGAAAAAGCAGAAGCTGATGCTAAACTGCTTGGAAACTCCCTGC
TGAAACTGGAAGGTGATGACCGTCTAGACATGAACTGCACGCTGCCCTTCAGTGACCAGGTAGTGTTG
GTGGGCACCGAGGAAGGGCTCTACGCCCTGAATGTCTTGAAAAACTCCCTAACCCATGTCCCAGGAAT
TGGAGCAGTCTTCCAAATTTATATTATCAAGGACCTGGAGAAGCTACTCATGATAGCAGGTGAAGAGC
GGGCACTGTGTCTTGTGGACGTGAAGAAAGTGAAACAGTCCCTGGCCCAGTCCCACCTGCCTGCCCAG
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GAACGGGCTCTGCATCTGTGCAGCCATTGTCTCCGCTACAACGAAAACCTCA
GCAAATACTGCATCCGGAAAGAGATAGAGACCTCAGAGCCCTGCAGCTGTATCCACTTCACCAATTAC
AGTATCCTCATTGGAACCAATAAATTCTACGAAATCGACATGAAGCAGTACACGCTCGAGGAATTCCT
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TCGTGCAGGTGAACAGCGCAGGGCAGCGAGAGGAGTACTTGCTGTGTTTCCACGAATTTGGAGTGTTC
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CTACAGAGAACCCTATCTGTTTGTGACCCACTTCAACTCACTCGAAGTAATTGAGATCCAGGCACGCT
CCTCAGCAGGGACCCCTGCCCGAGCGTACCTGGACATCCCGAACCCGCGCTACCTGGGCCCTGCCATT
TCCTCAGGAGCGATTTACTTGGCGTCCTCATACCAGGATAAATTAAGGGTCATTTGCTGCAAGGGAAA
CCTCGTGAAGGAGTCCGGCACTGAACACCACCGGGGCCCGTCCACCTCCCGCAGCAGCCCCAACAAGC
GAGGCCCACCCACGTACAACGAGCACATCACCAAGCGCGTGGCCTCCAGCCCAGCGCCGCCCGAAGGC
CCCAGCCACCCGCGAGAGCCAAGCACACCCCACCGCTACCGCGAGGGGCGGACCGAGCTGCGCAGGGA
CAAGTCTCCTGGCCGCCCCCTGGAGCGAGGAAGTCCCCCGGCCGGATGCTCAGCACGCGGAGAGAGC
GGTCCCCCGGGAGGCTGTTTGAAGACAGCAGCAGGGGCCGGCTGCCTGCGGGAGCCGTGAGGACCCCG
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A
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ORF Start: ATG at 1			ORF Stop: TAA at 6160

	SEQ ID NO: 2	2053 aa	MW at 234700.1kD
NOV1a, CG106764-01 Protein Sequence	MLKFKYGARNPLDAGAAEPIASRASRLNLFFQGKPPFMTQQQMSPLSREGILDALFVLFEECSQPALM KIKHVSNFVRKCSDTIAELQELQPSAKDFEVRSLVGCGHFAEVQVVREKATGDIYAMKVMKKKALLAQ EQVSFFEEERNILSRSTSPWIPQLQYAFQDKNHLYLVMEYQPGGDLLSLLNRYEDQLDENLIQFYLAE LILAVHSVHLMGYVHRDIKPENILVDRTGHIKLVDFGSAAKMNSNKVNAKLPIGTPDYMAPEVLTVMN GDGKGTYGLDCDWWSVGVIAYEMIYGRSPFAEGTSARTFNNIMNFQRFLKFPDDPKVSSDFLDLIQSL LCGQKERLKFEGLCCHPFFSKIDWNNIRNAPPPFVPTLKSDDDTSNFDEPEKNSWVSSSPCQLSPSGF SGEELPFVGFSYSKALGILGRSESVVSGLDSPAKTSSMEKKLLIKSKELQDSQDKCHKMEQEMTRLHR RVSEVEAVLSQKEVELKASETQRSLLEQDLATYITECSSLKRSLEQARMEVSQEDDKALQLLHDIREQ SRKLQEIKEQEYQAQVEEMRLMMNQLEEDLVSARRRSDLYESELRESRLAAEEFKRKATECQHKLLKA KDQGKPEVGEYAKLEKINAEQQLKIQELQEKLEKAVKASTEATELLQNIRQAKERAERELEKLQNRED SSEGIRRKLVEAEERRHSLENKVKRLETMERRENRLKDDIOTKSOOIOOMADKILELEEKHREAOVSA

99

| | QHLEVHLKQKEQHYEEKIKVLDNQIKKDLADKETLENMMQRHEEEAHEKGKILSEQKAMINAMDSKIR SLEQRIVELSEANKLAANSSLFTQRNMKAQEEMISELRQQKFYLETQAGKLEAQNRKLEEQLEKISHQ DHSDKNRLLELETRLREVSLEHEEQKLELKRQLTELQLSLQERESQLTALQAARAALESQLRQAKTEL EETTAEAEEEIQALTAHRDEIQRKFDALRNSCTVITDLEEQLNQLTEDNAELNNQNFYLSKQLDEASG ANDEIVQLRSEVDHLRREITEREMQLTSQKQTMEALKTTCTMLEEQVMDLEALNDELLEKERQWEAWR SVLGDEKSQFECRVRELQRMLDTEKQSRARADQRITESRQVVELAVKEHKAEILALQQALKEQKLKAE SLSDKLNDLEKKHAMLEMNARSLQQKLETERELKQRLLEEQAKLQQQMDLQKNHIFRLTQGLQEALDR ADLLKTERSDLEYQLENIQVLYSHEKVKMEGTISQQTKLIDFLQAKMDQPAKKKKVPLQYNELKLALE KEKARCAELEEALQKTRIELRSAREEAAHRKATDHPHPSTPATARQQIAMSAIVRSPEHQPSAMSLLA PPSSRRKESSTPEEFSRRLKERMHHNIPHRFNVGLNMRATKCAVCLDTVHFGRQASKCLECQVMCHPK CSTCLPATCGLPAEYATHFTEAFCRDKMNSPGLQTKEPSSSLHLEGWMKVPRNNKRGQQGWDRKYIVL EGSKVLIYDNEAREAGQRPVEEFELCLPDGDVSIHGAVGASELANTAKADVPYILKMESHPHTTCWPG RTLYLLAPSFPDKQRWVTALESVVAGGRVSREKAEADAKLLGNSLLKLEGDDRLDMNCTLPFSDQVVL VGTEEGLYALNVLKNSLTHVPGIGAVFQIYIIKDLEKLLMIAGEERALCLVDVKKVKQSLAQSHLPAQ PDISPNIFEAVKGCHLFGAGKIENGLCICAAMPSKVVILRYNENLSKYCIRKEIETSEPCSCIHFTNY SILIGTNKFYEIDMKQYTLEEFLDKNDHSLAPAVFAASSNSFPVSIVQVNSAGQREEYLLCFHEFGVF VDSYGRRSRTDDLKWSRLPLAFAYREPYLFVTHFNSLEVIEIQARSSAGTPARAYLDIPNPRYLGPAI SSGAIYLASSYQDKLRVICCKGNLVKESGTEHHRGPSTSRSSPNKRGPPTYNEHITKRVASSPAPPEG PSHPREPSTPHRYREGRTELRRDKSPGRPLEREKSPGRMLSTRRERSPGRLFEDSSRGRLPAGAVRTP LSQVNKVWDQSSV |

	SEQ ID NO: 3	1870 bp
NOV1b, 268667493 DNA Sequence	CACCGGTACCACCATGTTGAAGTTCAAATATGGAGCGCGGAATCCTTTGGATGCTGGTGCTGCTGAA CCCATTGCCAGCCGGGCCTCCAGGCTGAATCTGTTCTTCCAGGGGAAACCACCCTTTATGACTCAAC AGCAGATGTCTCCTCTTTCCCGAGAAGGGATATTAGATGCCCTCTTTGTTCTCTTTGAAGAATGCAG TCAGCCTGCTCTGATGAAGATTAAGCACGTGAGCAACTTTGTCCGGAAGTATTCCGACACCATAGCT GAGTTACAGGAGCTCCAGCCTTCGGCAAAGGACTTCGAAGTCAGAAGTCTTGTAGGTTGTGGTCACT TTGCTGAAGTGCAGGTGGTAAGAGAGAAAGCAACCGGGGACATCTATGCTATGAAAGTGATGAAGAA GAAGGCTTTATTGGCCCAGGAGCAGGTTTCATTTTTTGAGGAAGAGCGGAACATATTATCTCGAAGC ACAAGCCCGTGGATCCCCCAATTACAGTATGCCTTTCAGGACAAAAATCACCTTTATCTGGTCATGG AATATCAGCCTGGAGGGGACTTGCTGTCACTTTTGAATAGATATGAGGACCAGTTAGATGAAAACCT GATACAGTTTTACCTAGCTGAGCTGATTTTGGCTGTTCACAGCGTTCATCTGATGGGATACGTGCAT CGAGACATCAAGCCTGAGAACATTCTCGTTGACCGCCACAGGACACATCAAGCTGGTGGATTTTGGAT CTGCCGCGAAAATGAATTCAAACAAGATGGTGAATGCCAAACTCCCGATTGGGACCCCAGATTACAT GGCTCCTGAAGTGCTGACTGTGATGAACGGGGATGGAAAAGGCACCTACGGCCTGGACTGTGACTGG TGGTCAGTGGGCGTGATTGCCTATGAGATGATTTATGGGAGATCCCCCTTCGCAGAGGGAACCTCTG CCAGAACCTTCAATAACATTATGAATTTCCAGCGGTTTTTGAAATTTCCAGATGACCCCAAAGTGAG CAGTGACTTTCTTGATCTGATTCAAAGCTTGTTGTGCGGCCAGAAAGAGACTGAAGTTTGAAGGT CTTTGCTGCCATCCTTTCTTCTCTAAAATTGACTGGAACAACATTCGTAACTCTCCTCCCCCCTTCG TTCCCACCCTCAAGTCTGACGATGACACCTCCAATTTTGATGAACCAGAGAAGAATTCGTGGGTTTC ATCCTCTCCGTGCCAGCTGAGCCCCTCAGGCTTCTCGGGTGAAGAACTGCCGTTTGTGGGGTTTTCG TACAGCAAGGCACTGGGGATTCTTGGTAGATCTGAGTCTGTTGTGTCGGGTCTGGACTCCCCTGCCA AGACTAGCTCCATGGAAAAGAAACTTCTCATCAAAAGCAAAGAGCTACAAGACTCTCAGGACAAGTG TCACAAGATGGAGCAGGAAATGACCCGGTTACATCGGAGAGTGTCAGAGGTGGAGGCTGTGCTTAGT CAGAAGGAGGTGGAGCTGAAGGCCTCTGAGACTCAGAGATCCCTCCTGGGAGCAGGACCTTGCTACCT ACATCACAGAATGCAGTAGCTTAAAGCGAAGTTTGGAGCAAGCACGGATGGAGGTGTCCCAGGAGGA TGACAAAGCACTGCAGCTTCTCCATGATATCAGAGAGCAGAGCCGGAAGCTCCAAGAAATCAAAGAG CAGGAGTACCAGGCTCAAGTGGAAGAAATGAGGTTGATGATGAATCAGTTGGAAGAGGATCTTGTCT CAGCAAGAAGCAGCGGAGTGATCTCTACGAATCTGAGCTGAGAGAGTCTCGGCTTGCTGCTGAAGAATT CAAGCGGAAAGCGACAGAATGTCAGCATAAACTGTTGAAGGCTAAGGATCAGGTCGACGGC
	ORF Start: at 2	ORF Stop: end of sequence

	SEQ ID NO: 4	623 aa	MW at 70970.0kD
NOV1b, 268667493 Protein Sequence	TGTTMLKFKYGARNPLDAGAAEPIASRASRLNLFFQGKPPFMTQQQMSPLSREGILDALFVLFEECS QPALMKIKHVSNFVRKYSDTIAELQELQPSAKDFEVRSLVGCGHFAEVQVVREKATGDIYAMKVMKK KALLAQEQVSFFEEERNILSRSTSPWIPQLQYAFQDKNHLYLVMEYQPGGDLLSLLNRYEDQLDENL IQFYLAELILAVHSVHLMGYVHRDIKPENILVDRTGHIKLVDFGSAAKMNSNKMVNAKLPIGTPDYM APEVLTVMNGDGKGTYGLDCDWWSVGVIAYEMIYGRSPFAEGTSARTFNNIMNFQRFLKFPDDPKVS SDFLDLIQSLLCGQKERLKFEGLCCHPFFSKIDWNNIRNSPPPFVPTLKSDDDTSNFDEPEKNSWVS SSPCQLSPSGFSGEELPFVGFSYSKALGILGRSESVVSGLDSPAKTSSMEKKLLIKSKELQDSQDKC HKMEQEMTRLHRRVSEVEAVLSQKEVELKASETQRSLLEQDLATYITECSSLKRSLEQARMEVSQED

DKALQLLHDIREQSRKLQEIKEQEYQAQVEEMRLMMNQLEEDLVSARRRSDLYESELRESRLAAEEF
KRKATECQHKLLKAKDQVDG

	SEQ ID NO: 5	2497 bp
NOV1c, 268667539 DNA Sequence	CACCGGTACCCAGGGGAAGCCTGAAGTGGGAGAATATGCGAAACTGGAGAAGATCAATGCTGAGCAGC AGCTCAAAATTCAGGAGCTCCAAGAGAAACTGGAGAAGGCTGTAAAAGCCAGCACGGAGGCCACCGAG CTGCTGCAGAATATCCGCCAGGCAAAGGAGCGAGCCGAGAGGGAGCTGGAGAAGCTGCAGAACCGAGA GGATTCTTCTGAAGGCATCAGAAAGAAGCTGGTGGAAGCTGAGGAACGCCGCCATTCTCTGGAGAACA AGGTAAAGAGACTAGAGACCATGGAGCGTAGAGAAAACAGACTGAAGGATGACATCCAGACAAAATCC CAACAGATCCAGCAGATGGCTGATAAAATTCTGGAGCTCGAAGAGAAACATCGGGAGGCCCAAGTCTC AGCCCAGCACCTAGAAGTGCACCTGAAACAGAAAGAGCAGCACTATGAGGAAAAGATTAAAGTGTTGG ACAATCAGATAAAGAAAGACCTGGCTGACAAGGAGCACTGGAGAACATGATGCAGAGACACGAGGAG GAGGCCCATGAGAAGGGCAAAATTCTCAGCGAACAGAAGGCGATGATCAATGCTATGGATTCCAAGAT CAGATCCCTGGAACAGAGGATTGTGGAACTGTCTGAAGCCAATAAACTTGCAGCAAATAGCAGTCTTT TTACCCAAAGGAACATGAAGCCCAAGAAGATGATTTCTGAACTCAGGCAACAGAAATTTTACCTG GAGACACAGGCTGGGAAGTTGGAGGCCCAGAACCGAAAACTGGAGGAGCAGCTGGAGAAGATCAGCCA CCAAGACCACAGTGACAAGAATCGGCTGCTGGAACTGGAGACAAGATTGCGGGAGGTCAGTCTAGAGC ACGAGGAGCAGAAACTGGAGCTCAAGCGCCAGCTCACAGAGCTACAGCTCTCCCTGCAGGAGCGCGAG TCACAGTTGACAGCCCTGCAGGCTGCACGGGCCGGCCCTGGAGAGCCAGCTTCGCCAGGCGAAGACAGA GCTGGAAGAGACCAGCAGGAAGCTGAAGAGGAGGATCCAGGCACTCACGGCACATAGAGATGAAATCC AGCGCAAATTTGATGCTCTTCGTAACAGCTGTACTGTAATCACAGACCTGGAGGAGCAGCTAAACCAG CTGACCGAGGACAACGCTGAACTCAACAACCAAAACTTCTACTTGTCCAAACAACTCGATGAGGCTTC TGGCGCCAACGACGAGATTGTACAACTGCGAAGTGAAGTGGACCATCTCCGCCGGGAGATCACGGAAC GAGAGATGCAGCTTACCAGCCAGAAGCAAACGATGGAGGCTCTGAAGACCACGTGCACCATGCTGGAG GAACAGGTCATGGATTTGGAGGCCCTAAACGATGAGCTGCTAGAAAAAGAGCGGCAGTGGGAGGCCTG GAGGAGCGTCCTGGGTGATGAGAAATCCCAGTTTGAGTGTCGGGTTCGAGAGCTGCAGAGAATGCTGG ACACCGAGAAACAGAGCAGGGCGAGAGCCGATCAGCGGATCACCGAGTCTCGCCAGGTGGTGGAGCTG GCAGTGAAGGAGCACAAGGCTGAGATTCTCGCTCTGCAGCAGGCTCTCAAAGAGCAGAAGCTGAAGGC CGAGAGCCTCTCTGACAAGCTCAATGACCTGGAGAAGAAGCATGCTATGCTTGAAATGAATGCCCGAA GCTTACAGCAGAAGCTGGAGACTGAACGAGAGCTCAAACAGAGGCTTCTGGAAGAGCAAGCCAAATTA CAGCAGCAGATGGACCTGCAGAAAAATCACATTTTCCGTCTGACTCAAGGACTGCAAGAAGCTCTAGA TCGGGCTGATCTACTGAAGACAGAAAGAAGTGACTTGGAGTATCAGCTGGAAAACATTCAGGTTCTCT ATTCTCATGAAAAGGTGAAAATGGAAGGCACTATTTCTCAACAAACCAAACTCATTGATTTTCTGCAA GCCAAAATGGACCAACCTGCTAAAAAGAAAAAGGTTCCTCTGCAGTACAATGAGCTGAAGCTGGCCCT GGAGAAGGAGAAAGCTCGCTGTGCAGAGCTAGAGGAAGCCCTTCAGAAGACCCGCATCGAGCTCCGGT CCGCCCGGGAGGAAGCTGCCCACCGCAAAGCAACGGACCACCCACACCCATCCACGCCAGCCACCGCG AGGCAGCAGATCGCCATGTCCGCCATCGTGCGGTCGCCAGAGCACCAGCCCAGTGCCATGAGCCTGCT GGCCCCGCCATCCAGCCGCAGAAAGGAGTCTTCAACTCCAGAGGAATTTAGTCGGCGTCTTAAGGAAC GCATGCACCACAATATTCCTCACCGATTCAACGTAGGACTGAACATGCGAGCCACAAAGTGTGCTGTG TGTCTGGATACCGTGCACTTTGGACGCCAGGCATCCAAATGTCTCGAATGTCAGGTGATGTGTCACCC CAAGTGCTCCACGTGCTTGCCAGCCACCTGCGGCTTGCCTGTCGACGGC
	ORF Start: at 2	ORF Stop: end of sequence

	SEQ ID NO: 6	832 aa	MW at 96885.8kD
NOV1c, 268667539 Protein Sequence	TGTQGKPEVGEYAKLEKINAEQQLKIQELQEKLEKAVKASTEATELLQNIRQAKERAERELEKLQNRE DSSEGIRKKLVEAEERRHSLENKVKRLETMERRENRLKDDIQTKSQQIQQMADKILELEEKHREAQVS AQHLEVHLKQKEQHYEEKIKVLDNQIKKDLADKETLENMMQRHEEEAHEKGKILSEQKAMINAMDSKI RSLEQRIVELSEANKLAANSSLFTQRNMKAQEEMISELRQQKFYLETQAGKLEAQNRKLEEQLEKISH QDHSDKNRLLELETRLREVSLEHEEQKLELKRQLTELQLSLQERESQLTALQAARAALESQLRQAKTE LEETTAEAEEEIQALTAHRDEIQRKFDALRNSCTVITDLEEQLNQLTEDNAELNNQNFYLSKQLDEAS GANDEIVQLRSEVDHLRREITEREMQLTSQKQTMEALKTTCTMLEEQVMDLEALNDELLEKERQWEAW RSVLGDEKSQFECRVRELQRMLDTEKQSRARADQRITESRQVVELAVKEHKAEILALQQALKEQKLKA ESLSDKLNDLEKKHAMLEMNARSLQQKLETERELKQRLLEEQAKLQQQMDLQKNHIFRLTQGLQEALD RADLLKTERSDLEYQLENIQVLYSHEKVKMEGTISQQTKLIDFLQAKMDQPAKKKKVPLQYNELKLAL EKEKARCAELEEALQKTRIELRSAREEAAHRKATDHPHPSTPATARQQIAMSAIVRSPEHQPSAMSLL APPSSRRKESSTPEEFSRRLKERMHHNIPHRFNVGLNMRATKCAVCLDTVHFGRQASKCLECQVMCHP KCSTCLPATCGLPVDG

	SEQ ID NO: 7	2542 bp	
NOV1d, 268667543 DNA Sequence	CACCGGTACCCAGGGGAAGCCTGAAGTGGGAGAATATGCGAAACTGGAGAAGATCAATGCTGAGCAG CAGCTCAAAATTCAGGAGCTCCAAGAGAAACTGGAGAAGGCTGTAAAAGCCAGCACGGAGGCCACCG AGCTGCTGCAGAATATCCGCCAGGCAAAGGAGCGAGCCGAGAGGGAGCTGGAGAAGCTGCAGAACCG AGAGGATTCTTCTGAAGGCATCAGAAAGAAGCTGGTGGAAGCTGAGGAACGCCGCCATTCTCTGGAG AACAAGGTAAAGAGACTAGAGACCATGGAGCGTAGAGAAAACAGACTGAAGGATGACATCCAGACAA AATCCCAACAGATCCAGCAGATGGCTGATAAAATTCTGGAGCTCGAAGAGAAACATCGGGAGGCCCA AGTCTCAGCCCAGCACCTAGAAGTGCACCTGAAACAGAAAGAGCAGCACTATGAGGAAAAGATTAAA GTGTTGGACAATCAGATAAAGAAAGACCTGGCTGACAAGGAGACACTGGAGAACATGATGCAGAGAC ACGAGGAGGAGGCCCATGAGAAGGGCAAAATTCTCAGCGAACAGAAGGCGATGATCAATGCTATGGA TTCCAAGATCAGATCCCTGGAACAGAGGATTGTGGAACTGTCTGAAGCCAATAAACTTGCAGCAAAT AGCAGTCTTTTTACCCAAAGGAACATGAAGGCCCAAGAAGAGATGATTTCTGAACTCAGGCAACAGA AATTTTACCTGGAGACACAGGCTGGGAAGTTGGAGGCCCAGAACCGAAAACTGGAGGAGCAGCTGGA GAAGATCAGCCACCAAGACCACAGTGACAAGAATCGGCTGCTGGAACTGGAGACAAGATTGCGGGAG GTCAGTCTAGAGCACGAGGAGCAGAAACTGGAGCTCAAGCGCCAGCTCACAGAGCTACAGCTCTCCC TGCAGGAGCGCGAGTCACAGTTGACAGCCCTGCAGGCTGCACGGGCGGCCCTGGAGAGCCAGCTTCG CCAGGCGAAGACAGAGCTGGAAGAGACCACAGCAGAAGCTGAAGAGGAGATCCAGGCACTCACGGCA CATAGAGATGAAATCCAGCGCAAATTTGATGCTCTTCGTAACAGCTGTACTGTAATCACAGACCTGG AGGAGCAGCTAAACCAGCTGCACCGAGGACAACGCTGAACTCAACAACCAAAACTTCTACTTGTCCAA ACAACTCGATGAGGCTTCTGGCCGCCAACGACGAGATTGTACAACTGCGAAGTGAAGTGGACCATCTC CGCCGGGAGATCACGGAACGAGAGATGCAGCTTACCAGCCAGAAGCAAACGATGGAGGCTCTGAAGA CCACGTGCACCATGCTGGAGGAACAGGTCATGGATTTGGAGGCCCTAAACGATGAGCTGCTAGAAAA AGAGCGGCAGTGGGAGGCCTGGAGGAGCGTCCTGGGTGATGAGAAATCCCAGTTTGAGTGTCGGGTT CGAGAGCTGCAGAGGATGCTGGACACCGAGAAACAGAGCAGGCGGGCGGACCGATCAGCGGATCACCG AGTCTCGCCAGGTGGTGGAGCTGGCAGTGAAGGAGCACAAGGCTGAGATTCTCGCTCTGCAGCAGGC TCTCAAAGAGCAGAAGCTGAAGGCCGAGAGCCTCTCTGACAAGCTCAATGACCTGGAGAAGAAGCAT GCTATGCTTGAAATGAATGCCCGAAGCTTACAGCAGAAGCTGGAGACTGAACGAGAGCTCAAACAGA GGCTTCTGGAAGAGCAAGCCAAATTACAGCAGCAGATGGACCTGCAGAAAAATCACATTTTTCCGTCT GACTCAAGGACTGCAAGAAGCTCTAGATCGGGCTGATCTACTGAAGACAGAAAGAAGTGACTTGGAG TATCAGCTGGAAAACATTCAGGTTCTCTATTCTCATGAAAAGGTGAAAATGGAAGGCACTATTTCTC AACAAACCAAACTCATTGATTTTCTGCAAGCCAAAATGGACCAACCTGCTAAAAAGAAAAAGGGGTTT ATTTAGTCGACGGAAAGAGGACCCTGCTTTACCCACACAGGTTCCTCTGCAGTACAATGAGCTGAAG CTGGCCCTGGAAGAAGGAAAAGCTCGCTGTGCAGAGCTAGAGGAAGCCCTTCAGAAGACCCGCATCG AGCTCCGGTCCGCCCGGGAGGAAGCTGCCCACCGCAAAGCAACGGACCACCCACACCCATCCACGCC AGCCACCGCGAGGCAGCAGATCGCCATGTCTGCCATCGTGCGGTCGCCAGAGCACCAGCCCAGTGCC ATGAGCCTGCTGGCCCCGCCATCCAGCCGCAGAAAGGAGTCTTCAACTCCAGAGGAATTTAGTCGGC GTCTTAAGGAACGCATGCACCACAATATTCCTCACCGATTCAACGTAGGACTGAACATGCGAGCCAC AAAGTGTGCTGTGTGTCTGGATACCGTGCACTTTGGACGCCAGGCATCCAAATGTCTCGAATGTCAG GTGATGTGTCACCCCAAGTGCTCCACGTGCTTGCCAGCCACCTGCGGCTTGCCTGTCGACGGC		
	ORF Start: at 2		ORF Stop: end of sequence

5

	SEQ ID NO: 8	847 aa	MW at 98582.7kD
NOV1d, 268667543 Protein Sequence	TGTQGKPEVGEYAKLEKINAEQQLKIQELQEKLEKAVKASTEATELLQNIRQAKERAERELEKLQNR EDSSEGIRKKLVEAEERRHSLENKVKRLETMERRENRLKDDIQTKSQQIQQMADKILELEEKHREAQ VSAQHLEVHLKQKEQHYEEKIKVLDNQIKKDLADKETLENMMQRHEEEAHEKGKILSEQKAMINAMD SKIRSLEQRIVELSEANKLAANSSLFTQRNMKAQEEMISELRQQKFYLETQAGKLEAQNRKLEEQLE KISHQDHSDKNRLLELETRLREVSLEHEEQKLELKRQLTELQLSLQERESQLTALQAARAALESQLR QAKTELEETTAEAEEEIQALTAHRDEIQRKFDALRNSCTVITDLEEQLNQLTEDNAELNNQNFYLSK QLDEASGANDEIVQLRSEVDHLRREITEREMQLTSQKQTMEALKTTCTMLEEQVMDLEALNDELLEK ERQWEAWRSVLGDEKSQFECRVRELQRMLDTEKQSRARADQRITESRQVVELAVKEHKAEILALQQA LKEQKLKAESLSDKLNDLEKKHAMLEMNARSLQQKLETERELKQRLLEEQAKLQQQMDLQKNHIFRL TQGLQEALDRADLLKTERSDLEYQLENIQVLYSHEKVKMEGTISQQTKLIDFLQAKMDQPAKKKKGL FSRRKEDPALPTQVPLQYNELKLALEKEKARCAELEEALQKTRIELRSAREEAAHRKATDHPHPSTP ATARQQIAMSAIVRSPEHQPSAMSLLAPPSSRRKESSTPEEFSRRLKERMHHNIPHRFNVGLNMRAT KCAVCLDTVHFGRQASKCLECQVMCHPKCSTCLPATCGLPVDG		

10

	SEQ ID NO: 9	1870 bp	

| NOV1e,
268667555
DNA Sequence | CACCGGTACCTGCGGCTTGCCTGCTGAATATGCCACACACTTCACCGAGGCCTTCTGCCGTGACAAA
TGAACTCCCCAGGTCTCCAGACCAAGGAGCCCAGCAGCAGCTTGCACCTGGAAGGGTGGATGAAGGTG
CCCAGGAATAACAAACGAGGACAGCAAGGCTGGGACAGGAAGTACATTGTCCTGGAGGGATCAAAAGT
CCTCATTTATGACAATGAAGCCAGAGAAGCTGGACAGAGGCCGGTGGAAGAATTTGAGCTGTGCCTTC
CCGACGGGGATGTATCTATTCATGGTGCCGTTGGTGCTTCCGAACTCGCAAATACAGCCAAAGCAGAT
GTCCCATACATACTGAAGATGGAATCTCACCCGCACACCACCTGCTGGCCCGGGAGAACCCTCTACTT
GCTAGCTCCCAGCTTCCCTGACAAACAGCGCTGGGTCACCGCCTTAGAATCAGTTGTCGCAGGTGGGA
GAGTTTCTAGGGAAAAAGCAGAAGCTGATGCTAAACTGCTTGGAAACTCCCTGCTGAAACTGGAAGGT
GATGACCGTCTAGACATGAACTGCACGCTGCCCTTCAGTGACCAGGTGGTGTTGGTGGGTACCGAGGA
AGGGCTCTACGCCCTGAATGTCTTGAAAAACTCCCTAACCCATGTCCCAGGAATTGGAGCAGTCTTCC
AAATTTATATTATCAAGGACCTGGAGAAGCTACTCATGATAGCAGGAAGAGCGGGCACTGTGTCTT
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CAACATTTTTGAAGCTGTCAAGGGCTGCCACTTGTTTGGGGCAGGCAAGATTGAGAACGGGCTCTGCA
TCTGTGCAGCCATGCCCAGCAAGTCGTCATTCTCCGCTACAACGAAAACCTCAGCAAATACTGCATC
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GCCCCCTGGAGCGAGAGAAGTCCCCCGGCCGGATGCTCAGCACGCGGAGAGAGCGGTCCCCCGGGAGG
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CAAGGTGTGGGACCAGTCTTCAGTAGTCGACGGC |
| | ORF Start: at 2 | ORF Stop: end of sequence |

| | SEQ ID NO: 10 | 623 aa | MW at 69278.9kD |
| NOV1e,
268667555
Protein
Sequence | TGTCGLPAEYATHFTEAFCRDKMNSPGLQTKEPSSSLHLEGWMKVPRNNKRGQQGWDRKYIVLEGSKV
LIYDNEAREAGQRPVEEFELCLPDGDVSIHGAVGASELANTAKADVPYILKMESHPHTTCWPGRTLYL
LAPSFPDKQRWVTALESVVAGGRVSREKAEADAKLLGNSLLKLEGDDRLDMNCTLPFSDQVVLVGTEE
GLYALNVLKNSLTHVPGIGAVFQIYIIKDLEKLLMIAGEERALCLVDVKKVKQSLAQSHLPAQPDISP
NIFEAVKGCHLFGAGKIENGLCICAAMPSKVVILRYNENLSKYCIRKEIETSEPCSIHFTNYSILIG
TNKFYEIDMKQYTLEEFLDKNDHSLAPAVFAASSNSFPVSIVQVNSAGQREEYLLCFHEFGVFVDSYG
RRSRTDDLKWSRLPLAFAYREPYLFVTHFNSLEVIEIQARSSAGTPARAYLDIPNPRYLGPAISSGAI
YLASSYQDKLRVICCKGNLVKESGTEHHRGPSTSRSSPNKRGPPTYNEHITKRVASSPAPPEGPSHPR
EPSTPHRYREGRTELRRDKSPGRPLEREKSPGRMLSTRRERSPGRLFEDSSRGRLPAGAVRTPLSQVN
KVWDQSSVVDG |

| | SEQ ID NO: 11 | 1915 bp | |
| NOV1f,
268667574
DNA Sequence | CACCGGTACCTGCGGCTTGCCTGCTGAATATGCCACACACTTCACCGAGGCCTTCTGCCGTGATAAAA
TGAACTCCCCAGGTCTCCAGACCAAGGAGCCCAGCAGCAGCTTGCACCTGGAAGGGTGGATGAAGGTG
CCCAGGAATAACAAACGAGGACAGCAAGGCTGGGACAGGAAGTACATTGTCCTGGAGGGATCAAAAGT
CCTCATTTATGACAATGAAGCCAGAGAAGCTGGACAGAGGCCGGTGGAAGAATTTGAGCTGTGCCTTC
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GCTAGCTCCCAGCTTCCCTGACAAACAGCGCTGGGTCACCGCCTTAGAATCAGTTGTCGCAGGTGGGA
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TGGGTTCAGAAACTGCTTGGAAACTCCCTGCTGAAACTGGAAGGTGATGACCGTCTAGACATGAACTG
CACACTGCCCTTCAGTGACCAGGTGGTGTTGGTGGGCACCGAGGAAGGGCTCTACGCCCTGAATGTCT
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GTCCCTGGCCCAGTCCCACCTGCCTGCCCAGCCCGACATCTCACCCAACATTTTTGAAGCTGTCAAGG
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GTCGTCATTCTCCGCTACAACGAAAACCTCAGCAAATACTGCATCCGGAAAGAGATAGAGACCTCAGA
GCCCTGCAGCTGTATCCACTTCACCAATTACAGTATCCTCATTGGAACCAATAAATTCTACGAAATCG
ACATGAAGCAGTACACGCTCGAGGAATTCCTGGATAAGAATGACCATTCCTTGGCACCTGCTGTGTTT
GCCGCCTCTTCCAACAGCTTCCCTGTCTCAATCGTGCAGGTGAACAGCGCAGGGCAGCGAGAGGAGTA |

	CTTGCTGTGTTTCCACGAATTTGGAGTGTTCGTGGAT̶T̶C̶T̶A̶C̶G̶G̶A̶A̶G̶A̶C̶G̶T̶A̶C̶C̶G̶C̶A̶C̶A̶C̶A̶C̶A̶T̶C̶
	TCAAGTGGAGTCGCTTACCTTTGGCCTTTGCCTACAGAGAACCCTATCTGTTTGTGACCCACTTCAAC
	TCACTCGAAGTAATTGAGATCCAGGCACGCTCCTCAGCAGGGACCCCTGCCCGAGCGTACCTGGACAT
	CCCGAACCCGCGCTACCTGGGCCCTGCCATTTCCTCAGGAGCGATTTACTTGGCGTCCTCATACCAGG
	ATAAATTAAGGGTCATTTGCTGCAAGGGAAACCTCGTGAAGGAGTCCGGCACTGAACACCACCGGGGC
	CCGTCCACCTCCCGCAGCAGCCCCAACAAGCGAGGCCCACCCACGTACAACGAGCACATCACCAAGCG
	CGTGGCCTCCAGCCCAGCGCCGCCCGAAGGCCCCAGCCACCCGCGAGAGCCAAGCACACCCCACCGCT
	ACCGCGAGGGGCGGACCGAGCTGCGCAGGGACAAGTCTCCTGGCCGCCCCCTGGAGCGAGAGAAGTCC
	CCCGGCCGGATGCTCAGCACGCGGAGAGAGCGGTCCCCCGGGAGGCTGTTTGAAGACAGCAGCAGGGG
	CCGGCTGCCTGCGGGAGCCGTGAGGACCCCGCTGTCCCAGGTGAACAAGGTGTGGGACCAGTCTTCAG
	TAGTCGACGGC
ORF Start: at 2	ORF Stop: end of sequence

	SEQ ID NO: 12	638 aa	MW at 71010.8kD
NOV1f, 268667574 Protein Sequence	TGTCGLPAEYATHFTEAFCRDKMNSPGLQTKEPSSSLHLEGWMKVPRNNKRGQQGWDRKYIVLEGSKV LIYDNEAREAGQRPVEEFELCLPDGDVSIHGAVGASELANTAKADVPYILKMESHPHTTCWPGRTLYL LAPSFPDKQRWVTALESVVAGGRVSREKAEADAARDCVSYELLPAWVQKLLGNSLLKLEGDDRLDMNC TLPFSDQVVLVGTEEGLYALNVLKNSLTHVPGIGAVFQIYIIKDLEKLLMIAGEERALCLVDVKKVKQ SLAQSHLPAQPDISPNIFEAVKGCHLFGAGKIENGLCICAAMPSKVVILRYNENLSKYCIRKEIETSE PCSCIHFTNYSILIGTNKFYEIDMKQYTLEEFLDKNDHSLAPAVFAASSNSFPVSIVQVNSAGQREEY LLCFHEFGVFVDSYGRRSRTDDLKWSRLPLAFAYREPYLFVTHFNSLEVIEIQARSSAGTPARAYLDI PNPRYLGPAISSGAIYLASSYQDKLRVICCKGNLVKESGTEHHRGPSTSRSSPNKRGPPTYNEHITKR VASSPAPPEGPSHPREPSTPHRYREGRTELRRDKSPGRPLEREKSPGRMLSTRRERSPGRLFEDSSRG RLPAGAVRTPLSQVNKVWDQSSVVDG		

	SEQ ID NO: 13	6201 bp	
NOV1g, CG106764-02 DNA Sequence	ATGTTGAAGTTCAAATATGGAGCGCGGAATCCTTTGGATGCTGGTGCTGCTGAACCCATTGCCAGCC GGGCCTCCAGGCTGAATCTGTTCTTCCAGGGGAAACCACCCTTTATGACTCAACAGCAGATGTCTCC TCTTTCCCGAGAAGGGATATTAGATGCCCTCTTTGTTCTCTTTGAAGAATGCAGTCAGCCTGCTCTG ATGAAGATTAAGCACGTGAGCAACTTTGTCCGGAAGTGTTCCGACACCATAGCTGAGTTACAGGAGC TCCAGCCTTCGGCAAAGGACTTCGAAGTCAGAAGTCTTGTAGGTTGTGGTCACTTTGCTGAAGTGCA GGTGGTAAGAGAGAAAGCAACCGGGGACATCTATGCTATGAAAGTGATGAAGAAGAAGGCTTTATTG GCCCAGGAGCAGGTTTCATTTTTTGAGGAAGAGCGGAACATATTATCTCGAAGCACAAGCCCGTGGA TCCCCCAATTACAGTATGCCTTTCAGGACAAAAATCACCTTTATCTGGTGATGGAATATCAGCCTGG AGGGGACTTGCTGTCACTTTTGAATAGATATGAGGACCAGTTAGATGAAAAACCTGATACAGTTTTAC CTAGCTGAGCTGATTTTGGCTGTTCACAGCGTTCATCTGATGGGATACGTGCATCGGGACATCAAGC CTGAGAACATTCTCGTTGACCGCACAGGACACATCAAGCTGGTGGATTTTGGATCTGCCGCGAAAAT GAATTCAAACAAGGTGAATGCCAAACTCCCGATTGGGACCCCAGATTACATGGCTCCTGAAGTGCTG ACTGTGATGAACGGGGATGGAAAAGGCACCTACGGCCTGGACTGTGACTGGTGGTCAGTGGGCGTGA TTGCCTATGAGATGATTTATGGGAGATCCCCCTTCGCAGGGGAACCTCTGCCAGAACCTTCAATAA CATTATGAATTTCCAGCGGTTTTTTGAAATTTCCAGATGACCCCAAAGTGAGCAGTGACTTTCTTGAT CTGATTCAAAGCTTGTTGTGCGGCCAGAAAGAGAGACTGAAGTTTGAAGGTCTTTGCTGCCATCCTT TCTTCTCTAAAATTGACTGGAACAACATTCGTAACGCTCCTCCCCCCTTCGTTCCCACCCTCAAGTC TGACGATGACACCTCCAATTTTGATGAACCAGAGAAGAATTCGTGGGTTTCATCCTCTCCGTGCCAG CTGAGCCCCTCAGGCTTCTCGGGTGAAGAACTGCCGTTTGTGGGGTTTTCGTACAGCAAGGCACTGG GGATTCTTGGTAGATCTGAGTCTGTTGTGTCGGGTCTGGACTCCCTGCCAAGACTAGCTCCATGGA AAAGAAACTTCTCATCAAAAGCAAAGAGCTACAAGACTCTCAGGACAAGTGTCACAAGATGGAGCAG GAAATGACCCGGTTACATCGGAGAGTGTCAGGAGGCTGTGCTTAGTCAGAAGGAGGTGGAGC TGAAGGCCTCTGAGACTCAGAGATCCCTCCTGGAGCAGGACCTTGCTACCTACATCACAGAATGCAG TAGCTTAAAGCGAAGTTTGGAGCAAGCACGGATGGAGGTGTCCCAGGAGGATGACAAAGCACTGCAG CTTCTCCATGATATCAGAGAGCAGAGCCGGAAGCTCCAAGAAATCAAAGAGCAGGAGTACCAGGCTC AAGTGGAAGAAATGAGGTTGATGAATGAAATCAGTTGGAAGAGGATCTTGTCTCAGCAAGAAGACGGAG TGATCTCTACGAATCTGAGCTGAGAGAGTCTCGGCTTGCTGCTGAAGAATTCAAGCGGAAAGCGACA GAATGTCAGCATAAACTGTTGAAGGCTAAGGATCAGGGGAAGCCTGAAGTGGGAGAATATGCGAAAC TGGAGAAGATCAATGCTGAGCAGCAGCTCAAAATTCAGGAGCTCCAAGAGAAACTGGAGAAGGCTGT AAAAGCCAGCACGGAGGCCCACCGAGCTGCTGCAGAATATCCGCCAGGCAAAGGAGCGAGCCGAGAGG GAGCTGGAGAAGCTGCAGAACCGAGAGGATTCTTCTGAAGGCATCAGAAAGAAGCTGGTGGAAGCTG AGGAACGCCGCCATTCTCTGGAACAAGGTAAAGAGACTAGAGACCATGGAGCGTAGAGAAAACAG ACTGAAGGATGACATCCAGACAAAATCCCAACAGATCCAGCAGATGGCTGATAAAATTCTGGAGCTC GAAGAGAAACATCGGGAGGCCCAAGTCTCAGCCCAGCACCTAGAAGTGCACCTGAAACAGAAAGAGC AGCACTATGAGGAAAAGATTAAAGTATTGGACAATCAGATAAAGAAAGACCTGGCTGACAAGGAGAC		

ACTGGAGAACATGATGCGAGAGACACGAGGAGGAGGCCCATGAGAAGGGCAAAATTCTCAGCGAACAG
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GCGGCCCTGGAGAGCCAGCTTCGCCAGGCGAAGACAGAGCTGGAAGAGACCACAGCAGAAGCTGAAG
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GCAAACGATGGAGGCTCTGAAGACCACGTGCACCATGCTGGAGGAACAGGTCATGGATTTGGAGGCC
CTAAACGATGAGCTGCTAGAAAAAGAGCGGCAGTGGGAGGCCTGGAGGAGCGTCCTGGGTGATGAGA
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TCAATGACCTGGAGAAGAAGCATGCTATGCTTGAAATGAATGCCCGAAGCTTACAGCAGAAGCTGGA
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CTGCCCAGCCCGACATCTCACCCAACATTTTTGAAGCTGTCAAGGGCTGCCACTTGTTTGGGGCAGG
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CGAGGAATTCCTGGATAAGAATGACCATTCCTTGGCACCTGCTGTGTTTGCCGCCTCTTCCAACAGC
TTCCCTGTCTCAATCGTGCAGGTGAACAGCGCAGGGCAGCGAGAGGAGTACTTGCTGTGTTTCCACG
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GAGATCCAGGCACTGCTCCTCAGCAGGGACCCCTGCCCGAGCGTACCTGGACATCCCGAACCCGCGCT
ACCTGGGCCCTGCCATTTCCTCAGGAGCGATTTACTTGGCGTCCTCATACCAGGATAAATTAAGGGT
CATTTGCTGCAAGGGGAAACCTCGTGAAGGAGTCCGGCACTGAACACCACCGGGGCCCGTCCACCTCC
CGCAGCAGCCCCAACAAGCGAGGCCCACCCACGTACAACGAGCACATCACCAAGCGCGTGGCCTCCA
GCCCAGCGCCGCCCGAAGGCCCCAGCCACCCGCGAGAGCCAAGCACACCCCACCGCTACCGCGAGGG
GCGGACCGAGCTGCGCAGGGACAAGTCTCCTGGCCGCCCCCCTGGAGCGAGAGAAGTCCCCCGGCCGG
ATGCTCAGCACGCGGAGAGAGCGGTCCCCCGGGAGGCTGTTTGAAGACAGCAGCAGGGGCCGGCTGC
CTGCGGGAGCCGTGAGGACCCCGCTGTCCCAGGTGAACAAGGTGAGGCAGCATTCCGAGGCCTGTGT
GTCTGTTGCGGAGGCCAGGAGTGACTTGGGGAACTGA

ORF Start: ATG at 1		ORF Stop: TGA at 6199

	SEQ ID NO: 14	2066 aa	MW at 236008.5kD
NOV1g, CG106764-02 Protein Sequence	MLKFKYGARNPLDAGAAEPIASRASRLNLFFQGKPPFMTQQQMSPLSREGILDALFVLFEECSQPAL MKIKHVSNFVRKCSDTIAELQELQPSAKDFEVRSLVGCGHFAEVQVVREKATGDIYAMKVMKKKALL AQEQVSFFEEEERNILSRSTSPWIPQLQYAFQDKNHLYLVMEYQPGGDLLSLLNRYEDQLDENLIQFY LAELILAVHSVHLMGYVHRDIKPENILVDRTGHIKLVDFGSAAKMNSNKVNAKLPIGTPDYMAPEVL TVMNGDGKGTYGLDCDWWSVGVIAYEMIYGRSPFAEGTSARTFNNIMNFQRFLKPPDDPKVSSDFLD LIQSLLCGQKERLKFEGLCCHPFFSKIDWNNIRNAPPPFVPTLKSDDDTSNFDEPEKNSWVSSSPCQ LSPSGFSGEELPFVGFSYSKALGILGRSESVVSGLDSPAKTSSMEKKLLIKSKELODSODKCHKMEO		

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EMTRLHRRVSEVEAVLSQKEVELKASETQRSLLEQDLATYITECSSLKRSLEQARMEVSQEDDKALQ
LLHDIREQSRKLQEIKEQEYQAQVEEMRLMMNQLEEDLVSARRRSDLYESELRESRLAAEEFKRKAT
ECQHKLLKAKDQGKPEVGEYAKLEKINAEQQLKIQELQEKLEKAVKASTEATELLQNIRQAKERAER
ELEKLQNREDSSEGIRKKLVEAEERRHSLENKVKRLETMERRENRLKDDIQTKSQQIQQMADKILEL
EEKHREAQVSAQHLEVHLKQKEQHYEEKIKVLDNQIKKDLADKETLENMMQRHEEEAHEKGKILSEQ
KAMINAMDSKIRSLEQRIVELSEANKLAANSSLFTQRNMKAQEEMISELRQQKFYLETQAGKLEAQN
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AALESQLRQAKTELEETTAEAEEEIQALTAHRDEIQRKFDALRNSCTVITDLEEQLNQLTEDNAELN
NQNFYLSKQLDEASGANDEIVQLRSEVDHLRREITEREMQLTSQKQTMEALKTTCTMLEEQVMDLEA
LNDELLEKERQWEAWRSVLGDEKSQFECRVRELQRMLDTEKQSRARADQRITESRQVVELAVKEHKA
EILALQQALKEQKLKAESLSDKLNDLEKKHAMLEMNARSLQQKLETERELKQRLLEEQAKLQQQMDL
QKNHIFRLTQGLQEALDRADLLKTERSDLEYQLENIQVLYSHEKVKMEGTISQQTKLIDFLQAKMDQ
PAKKKKVPLQYNELKLALEKEKARCAELEEALQKTRIELRSAREEAAHRKATDHPHPSTPATARQQI
AMSAIVRSPEHQPSAMSLLAPPSSRRKESSTPEEFSRRLKERMHHNIPHRFNVGLNMRATKCAVCLD
TVHFGRQASKCLECQVMCHPKCSTCLPATCGLPAEYATHFTEAFCRDKMNSPGLQTKEPSSSLHLEG
WMKVPRNNKRGQQGWDRKYIVLEGSKVLIYDNEAREAGQRPVEEFELCLPDGDVSIHGAVGASELAN
TAKADVPYILKMESHPHTTCWPGRTLYLLAPSFPDKQRWVTALESVVAGGRVSREKAEADAKLLGNS
LLKLEGDDRLDMNCTLPFSDQVVLVGTEEGLYALNVLKNSLTHVPGIGAVFQIYIIKDLEKLLMIAG
EERALCLVDVKKVKQSLAQSHLPAQPDISPNIFEAVKGCHLFGAGKIENGLCICAAMPSKVVILRYN
ENLSKYCIRKEIETSEPCSCIHFTNYSILIGTNKFYEIDMKQYTLEEFLDKNDHSLAPAVFAASSNS
FPVSIVQVNSAGQREEYLLCFHEFGVFVDSYGRRSRTDDLKWSRLPLAFAYREPYLFVTHFNSLEVI
EIQARSSAGTPARAYLDIPNPRYLGPAISSGAIYLASSYQDKLRVICCKGNLVKESGTEHHRGPSTS
RSSPNKRGPPTYNEHITKRVASSPAPPEGPSHPREPSTPHRYREGRTELRRDKSPGRPLEREKSPGR
MLSTRRERSPGRLFEDSSRGRLPAGAVRTPLSQVNKVRQHSEACVSVAEARSDLGN
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Sequence comparison of the above protein sequences yields the following sequence relationships shown in Table 1B.

5

Table 1B. Comparison of NOV1a against NOV1b through NOV1g.		
Protein Sequence	NOV1a Residues/ Match Residues	Identities/ Similarities for the Matched Region
NOV1b	1..615 5..620	601/616 (97%) 602/616 (97%)
NOV1c	615..1442 4..831	690/828 (83%) 691/828 (83%)
NOV1d	615..1442 4..846	690/843 (81%) 691/843 (81%)
NOV1e	1436..2053 3..620	618/618 (100%) 618/618 (100%)
NOV1f	1436..2053 3..635	618/633 (97%) 618/633 (97%)
NOV1g	1..2051 1..2051	1900/2051 (92%) 1900/2051 (92%)

Further analysis of the NOV1a protein yielded the following properties shown in

10   Table 1C.

Table 1C. Protein Sequence Properties NOV1a	
PSort analysis:	0.9800 probability located in nucleus; 0.3000 probability located in microbody (peroxisome); 0.1000 probability located in mitochondrial matrix space; 0.1000 probability located in lysosome (lumen)
SignalP analysis:	No Known Signal Sequence Predicted

5   A search of the NOV1a protein against the Geneseq database, a proprietary

database that contains sequences published in patents and patent publication, yielded

several homologous proteins shown in Table 1D.

Table 1D. Geneseq Results for NOV1a				
Geneseq Identifier	Protein/Organism/Length [Patent #, Date]	NOV1a Residues/ Match Residues	Identities/ Similarities for the Matched Region	Expect Value
AAU03501	Human protein kinase #1 - Homo sapiens, 2053 aa. [WO200138503-A2, 31-MAY-2001]	1..2051 1..2053	2044/2053 (99%) 2046/2053 (99%)	0.0
AAB43359	Human ORFX ORF3123 polypeptide sequence SEQ ID NO:6246 - Homo sapiens, 1286 aa. [WO200058473-A2, 05-OCT-2000]	768..2053 1..1286	1286/1286 (100%) 1286/1286 (100%)	0.0
ABB11117	Human RHO/RAC effector homologue, SEQ ID NO:1487 - Homo sapiens, 999 aa. [WO200157188-A2, 09-AUG-2001]	968..1947 1..980	976/980 (99%) 976/980 (99%)	0.0
AAU31443	Novel human secreted protein #1934 - Homo sapiens, 910 aa. [WO200179449-A2, 25-OCT-2001]	1114..1982 1..869	867/869 (99%) 867/869 (99%)	0.0
AAE16261	Human kinase PKIN-7 protein - Homo sapiens, 497 aa. [WO200196547-A2, 20-DEC-2001]	1..467 1..468	463/468 (98%) 465/468 (98%)	0.0

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In a BLAST search of public sequence datbases, the NOV1a protein was found to
have homology to the proteins shown in the BLASTP data in Table 1E.

5

Table 1E. Public BLASTP Results for NOV1a				
Protein Accession Number	Protein/Organism/Length	NOV1a Residues/ Match Residues	Identities/ Similarities for the Matched Portion	Expect Value
O88938	Rho/rac-interacting citron kinase - Mus musculus (Mouse), 2055 aa.	1..2053 1..2055	1974/2055 (96%) 2014/2055 (97%)	0.0
O88528	Citron-K kinase - Mus musculus (Mouse), 1641 aa (fragment).	373..2053 1..1641	1599/1683 (95%) 1616/1683 (96%)	0.0
P49025	Citron protein (Rho-interacting, serine/threonine kinase 21) - Mus musculus (Mouse), 1597 aa.	467..2053 9..1597	1563/1589 (98%) 1578/1589 (98%)	0.0
Q9QX19	Postsynaptic density protein - Rattus norvegicus (Rat), 1618 aa.	467..2053 1..1618	1556/1619 (96%) 1573/1619 (97%)	0.0
O14578	Citron protein (Rho-interacting, serine/threonine kinase 21) - Homo sapiens (Human), 1286 aa (fragment).	768..2053 1..1286	1286/1286 (100%) 1286/1286 (100%)	0.0

PFam analysis predicts that the NOV1a protein contains the domains shown in the
Table 1F.

10

Table 1F. Domain Analysis of NOV1a			
Pfam Domain	NOV1a Match Region	Identities/ Similarities for the Matched Region	Expect Value
pkinase	97..359	89/302 (29%) 196/302 (65%)	2.7e-62
pkinase_C	360..389	15/32 (47%) 24/32 (75%)	0.00023

DAG_PE-bind	1389..1437	14/51 (27%) 34/51 (67%)	6.1e-05
PH	1470..1589	20/121 (17%) 87/121 (72%)	1.8e-11
CNH	1618..1915	107/378 (28%) 289/378 (76%)	1.5e-110

Example 2.

The NOV2 clone was analyzed, and the nucleotide and encoded polypeptide
sequences are shown in Table 2A.

Table 2A. NOV2 Sequence Analysis			
	SEQ ID NO: 15	1238 bp	
NOV2a, CG117662-01 DNA Sequence	ATGGATGGATGGAGAAGGATGCCTCGCTGGGGACTGCTGCTGCTGCTCTGGGGCTCCTGTACCTTTGG TCTCCCGACAGACACCACCACCTTTAAACGGATCTTCCTCAAGAGAATGCCCTCAATCCGAGAAAGCC TGAAGGAACGAGGTGTGGACATGGCCAGGCTTGGTCCCGAGTGGAGCCAACCCATGAAGAGGCTGACA CTTGGCAACACCACCTCCTCCGTGATCCTCACCAACTACATGGACACCCAGTACTATGGCGAGATTGG CATCGGCACCCCACCCCAGACCTTCAAAGTCGTCTTTGACACTGGTTCGTCCAATGTTTGGGTGCCCT CCTCCAAGTGCAGCCGTCTCTACACTGCCTGTGTGTATCACAAGCTCTTCGATGCTTCGGATTCCTCC AGCTACAAGCACAATGGAACAGAACTCACCCTCCGCTATTCAACAGGGACAGTCAGTGGCTTTCTCAG CCAGGACATCATCACCGTGGGTGGAATCACGGTGACACAGATGTTTGGAGAGGTCACGGAGATGCCCG CCTTACCCTTCATGCTGGCCGAGTTTGATGGGGTTGTGGGCATGGGCTTCATTGAACAGGCCATTGGC AGGGTCACCCCTATCTTCGACAACATCATCTCCCAAGGGGTGCTAAAAGAGGACGTCTTCTCTTTCTA CTACAACAGAGATTCCGAGAATTCCCAATCGCTGGGAGGACAGATTGTGCTGGGGAGGCAGCGACCCCC AGCATTACGAAGGGAATTTCCACTATATCAACCTCATCAAGACTGGTGTCTGGCAGATTCAAATGAAG GGGGTGTCTGTGGGGTCATCCACCTTGCTCTGTGAAGACGGCTGCCTGGCATTGGTAGACACCGGTGC ATCCTACATCTCAGGTTCTACCAGCTCCATAGAGAAGCTCATGGAGGCCTTGGGAGCCAAGAAGAGGC TGTTTGATTATGTCGTGAAGTGTAACGAGGGCCCTACACTCCCCGACATCTCTTTCCACCTGGGAGGC AAAGAATACACGCTCACCAGCGCGGACTATGTATTTCAGGAATCCTACAGTAGTAAAAAGCTGTGCAC ACTGGCCATCCACGCCATGGATATCCCGCCACCCACTGGACCCACCTGGGCCCTGGGGGCCACCTTCA TCCGAAAGTTCTACACAGAGTTTGATCGGCGTAACAACCGCATTGGCTTCGCCTCGGCCCGCTGAGGC CCTCTGCCACCCAG		
	ORF Start: ATG at 1		ORF Stop: TGA at 1219

	SEQ ID NO: 16	406 aa	MW at 45030.9kD
NOV2a, CG117662-01 Protein Sequence	MDGWRRMPRWGLLLLLWGSCTFGLPTDTTTFKRIFLKRMPSIRESLKERGVDMARLGPEWSQPMKRLT LGNTTSSVILTNYMDTQYYGEIGIGTPPQTFKVVFDTGSSNVWVPSSKCSRLYTACVYHKLFDASDSS SYKHNGTELTLRYSTGTVSGFLSQDIITVGGITVTQMFGEVTEMPALPFMLAEFDGVVGMGFIEQAIG RVTPIFDNIISQGVLKEDVFSFYYNRDSENSQSLGGQIVLGGSDPQHYEGNFHYINLIKTGVWQIQMK GVSVGSSTLLCEDGCLALVDTGASYISGSTSSIEKLMEALGAKKRLFDYVVKCNEGPTLPDISFHLGG KEYTLTSADYVFQESYSSKKLCTLAIHAMDIPPPTGPTWALGATFIRKFYTEFDRRNNRIGFASAR		

	SEQ ID NO: 17	911 bp	
NOV2b, CG117662-02 DNA Sequence	ATGGATGGATGGAGAAGGATGCCTCGCTGGGGACTGCTGCTGCTGCTCTGGGGCTCCTGTACCTTTG GTCTCCCGACAGACACCACCACCTTTAAACGGATCTTCCTCAAGAGAATGCCCTCAATCCGAGAAAG CCTGAAGGAACGAGGTGTGGACATGGCCAGGCTTGGTCCCGAGTGGAGCCAACCCATGAAGAGGCTG ACACTTGGCAACACCACCTCCTCCGTGATCCTCACCAACTACATGGACACCCAGTACTATGGCGAGA TTGGCATCGGCACCCCACCCCAGACCTTCAAAGTCGTCTTTGACACTGGTTCGTCCAATGTTTGGGT GCCCTCCTCCAAGTGCAGCCGTCTCTACACTGCCTGTGTGTATCACAAGCTCTTCGATGCTTCGGAT TCCTCCAGCTACAAGCACAATGGAACAGAACTCACCCTCCGCTATTCAACAGGGACAGTCAGTGGCT TTCTCAGCCAGGACATCATCACCGTGTCTGTGGGGTCATCCACCTTACTCTGTGAAGACGGCTGCCT GGCATTGGTAGACACCGGTGCATCCTACATCTCAGGTTCTACCAGCTCCATAGAGAAGCTCATGGAG GCCTTGGGAGCCAAGAAGAGGCTGTTTGATTATGTCGTGAAGTGTAACGAGGGCCCTACACTCCCCG ACATCTCTTTCCACCTGGGAGGCAAAGAATACACGCTCACCAGCGCGGACTATGTATTTCAGGAATC CTACAGTAGTAAAAAGCTGTGCACACTGGCCATCCACGCCATGGATATCCCGCCACCCACTGGACCC ACCTGGGCCCTGGGGGCCACCTTCATCCGAAAGTTCTACACAGAGTTTGATCGGCGTAACAACCGCA TTGGCTTCGCCTCGGCCCGCTGAGGCCCTCTGCCACCCAG		
	ORF Start: ATG at 1		ORF Stop: TGA at 892

	SEQ ID NO: 18	297 aa	MW at 33025.3kD
NOV2b, CG117662-02 Protein Sequence	MDGWRRMPRWGLLLLLWGSCTFGLPTDTTTFKRIFLKRMPSIRESLKERGVDMARLGPEWSQPMKRL TLGNTTSSVILTNYMDTQYYGEIGIGTPPQTFKVVFDTGSSNVWVPSSKCSRLYTACVYHKLFDASD SSSYKHNGTELTLRYSTGTVSGFLSQDIITVSVGSSTLLCEDGCLALVDTGASYISGSTSSIEKLME ALGAKKRLFDYVVKCNEGPTLPDISFHLGGKEYTLTSADYVFQESYSSKKLCTLAIHAMDIPPPTGP TWALGATFIRKFYTEFDRRNNRIGFASAR		

Sequence comparison of the above protein sequences yields the following sequence relationships shown in Table 2B.

Table 2B. Comparison of NOV2a against NOV2b.		
Protein Sequence	NOV2a Residues/ Match Residues	Identities/ Similarities for the Matched Region
NOV2b	1..165 1..165	165/165 (100%) 165/165 (100%)

Further analysis of the NOV2a protein yielded the following properties shown in Table 2C.

Table 2C. Protein Sequence Properties NOV2a	
PSort analysis:	0.3700 probability located in outside; 0.2541 probability located in microbody (peroxisome); 0.1900 probability located in lysosome (lumen); 0.1000 probability located in endoplasmic reticulum (membrane)
SignalP analysis:	Cleavage site between residues 24 and 25

A search of the NOV2a protein against the Geneseq database, a proprietary database that contains sequences published in patents and patent publication, yielded several homologous proteins shown in Table 2D.

Table 2D. Geneseq Results for NOV2a				
Geneseq Identifier	Protein/Organism/Length [Patent #, Date]	NOV2a Residues/ Match Residues	Identities/ Similarities for the Matched Region	Expect Value
AAW23244	Human renin - Homo sapiens, 406 aa. [WO9728684-A1, 14-AUG-1997]	1..406 1..406	404/406 (99%) 404/406 (99%)	0.0
AAP50135	Sequence of pre-pro-renin - Homo sapiens, 406 aa. [EP135347-A, 27-MAR-1985]	1..406 1..406	404/406 (99%) 404/406 (99%)	0.0
ABB11781	Human renin homologue, SEQ ID NO:2151 - Homo sapiens, 438 aa. [WO200157188-A2, 09-AUG-2001]	1..406 31..438	391/408 (95%) 393/408 (95%)	0.0
AAU72879	Human aspartyl protease partial protein sequence #4 - Homo sapiens, 412 aa. [WO200183782-A2, 08-NOV-2001]	24..405 14..409	169/400 (42%) 246/400 (61%)	1e-90
AAY93685	Amino acid sequence of novel polypeptide PRO292 - Homo sapiens, 412 aa. [WO200037640-A2, 29-JUN-2000]	24..405 14..409	169/400 (42%) 246/400 (61%)	1e-90

In a BLAST search of public sequence datbases, the NOV2a protein was found to have homology to the proteins shown in the BLASTP data in Table 2E.

Table 2E. Public BLASTP Results for NOV2a

Protein Accession Number	Protein/Organism/Length	NOV2a Residues/ Match Residues	Identities/ Similarities for the Matched Portion	Expect Value
P00797	Renin precursor, renal (EC 3.4.23.15) (Angiotensinogenase) - Homo sapiens (Human), 406 aa.	1..406 1..406	405/406 (99%) 405/406 (99%)	0.0
Q9TSZ1	Preprorenin precursor (EC 3.4.23.15) - Callithrix jacchus (Common marmoset), 400 aa.	1..406 1..400	381/406 (93%) 389/406 (94%)	0.0
P52115	Renin precursor, renal (EC 3.4.23.15) (Angiotensinogenase) - Ovis aries (Sheep), 400 aa.	7..406 1..400	292/401 (72%) 338/401 (83%)	e-175
Q15296	Kidney mRNA fragment for renin (Aa 105-401) - Homo sapiens (Human), 300 aa (fragment).	108..406 1..300	297/300 (99%) 298/300 (99%)	e-172
P06281	Renin precursor, renal (EC 3.4.23.15) (Angiotensinogenase) - Mus musculus (Mouse), 402 aa.	5..406 4..402	281/403 (69%) 331/403 (81%)	e-167

PFam analysis predicts that the NOV2a protein contains the domains shown in the Table 2F.

5

Table 2F. Domain Analysis of NOV2a			
Pfam Domain	NOV2a Match Region	Identities/ Similarities for the Matched Region	Expect Value
asp	31..405	174/428 (41%) 339/428 (79%)	4.1e-197

Example 3.

10   The NOV3 clone was analyzed, and the nucleotide and encoded polypeptide sequences are shown in Table 3A.

Table 3A. NOV3 Sequence Analysis

	SEQ ID NO: 19	2827 bp	
NOV3a, CG118051-01 DNA Sequence	TGGCGATGCTACTGTTTAATTGCAGGAGGTGGGGGTGTGTGTACCATGTACCAGGGCTATTAGAAGCA AGAAGGAAGGAGGGAGGGCAGAGCGCCCTGCTGAGCAACAAAGGACTCCTGCAGCCTTCTCTGTCTGT CTCTTGGCACAGGCACATGGGGAGGCCTCCCGCAGGTGGGGGGGCCACCAGTCCAGGGGTGGGAGCACT ACAGGGCACGAGTTGGTTTGGGAGCTGCCAGTCTCCTGGGAGGATCGCAGTCAGCAGAGCAGGGCTGA GGCCTGGGGGTAGGAGCAGAGCCTGCGCATCTGGAGGCAGCATGTCCAAGAAAGGGAGTGGAGGTGCA GCGAAGGACCCAGGGGCAGAGCCCACGCTGGGGATGGACCCCTTCGAGGACACACTGCGGCGGCTGCG TGAGGCCTTCAACTGAGGGCGCACGCGGCCGGCCGAGTTCCGGGCTGCGCAGCTCCAGGGCCTGGGCC ACTTCCTTCAAGAAAACAAGCAGCTTCTGCGCGACGTGCTGGCCCAGGACCTGCATAAGCCAGCTTTC GAGGCAGACATATCTGAGCTCATCCTTTGCCAGAACGAGGTTGACTACGCTCTCAAGAACCTTCAGGC CTGGATGAAGGATGAACCACGGTCCACGAACCTGTTCATGAAGCTGGACTCGGTCTTCATCTGGAAGG AACCCTTTGGCCTGGTCCTCATCATCGCACCCTGGAACTACCCATTGAACCTGACCCTGGTGCTCCTG GTGGGCACCCTCCCCGCAGGGAATTGCGTGGTGCTGAAGCCGTCAGAAATCAGCCAGGGCACAGAGAA GGTCCTGGCTGAGGTGCTGCCCCCAGTACCTGGACCAGGACCTGCTTTGCCGTGGTGCTGGGCGGACCCC AGGAGACAGGGCAGCTGCTAGAGCACAAGTTGGACTACATCTTCTTCACAGGGAGCCCTCGTGTGGGC AAGATTGTCATGACTGCTGCCACCAAGCACCTGACGCCTGTCACCCTGGAGCTGGGGGGGCAAGAACCC CTGCTACGTGGACGACAACTGCGACCCCCAGACCGTGGCCAACCGCGTGGCCTGGTTCTGCTACTTCA ATGCCGGCCAGACCTGCGTGGCCCCTGACTACGTCCTGTGCAGCCCCGAGATGCAGGAGAGGCTGCTG CCCGCCCTGCAGAGCACCATCACCCGTTTCTATGGCGACGACCCCCAGTCCTCCCCAAACCTGGGCCG CATCATCAACCAGAAACAGTTCCAGCGGCTGCGGGCATTGCTGGGCTGCGGCCGCGTGGCCATTGGGG GCCAGAGCAACGAGAGCGATCGCTACATCGCCCCCACGGTGCTGGTGGACGTGCAGGAGACGGAGCCT GTGATGCAGGAGGAGATCTTCGGGCCCATCCTGCCCATCGTGAACGTGCAGAGCGTGGACGAGGCCAT CAAGTTCATCAACCGGCAGGAGAAGCCCCTGGCCCTGTACGCCTTCTCCAACAGCAGACAGGTTGTGA ACCAGATGCTGGAGCGGACCAGCAGCGGCAGCTTTGGAGGCAATGAGGGCTTCACCTACATATCTCTG CTGTCCGTGCCATTCGGGGGAGTCGGCCACAGTGGGATGGGCCGGTACCACGGCAAGTTCACCTTCGA CACCTTCTCCCACCACCGCACCTGCCTGCTCGCCCCCTCCGGCCTGGAGAAATTAAAGGAGATCCGCT ACCCACCCTATACCGACTGGAACCAGCAGCTGTTACGCTGGGGCATGGGCTCCCAGAGCTGCACCCTC CTGTGAGCGTCCCACCCGCCTCCAACGGGTCACACAGAGAAACCTGAGTCTAGCCATGAGGGGCTTAT GCTCCCAACTCACATTGTTCCTCCAGACCGCAGGCTCCCCCAGCCTCAGGTTGCTGGAGCTGTCACAT GACTGCATCCTGCCTGCCAGGGCTGCAAAGCAAGGTCTTGCTTCTATCTGGGGGACGCTGCTCGAGAG AGGCCGAGAGGCCGACAGAACATGCCAGGTGTCCTCACTCACCCCACCCTCCCCAATTCCAGCCCTTTG CCCTCTCGGTCAGGGTTGGCCAGGCCCAGTCACAGGGGCAGTGTCACCCTGGAAAATACAGTGCCCTG CCTTCTTAGGGGCATCAGCCCTGAACGGTTGAGAGCGTGGAGCCCTCCAGGCCTTTGCTCTCCCCTCT AGGCACACGCGCACTTCCACCTCTGCCCCATCCCAACTGCACCAGCACTGCCTCCCCCAGGGATCCTC TCACATCCCACACTGGTCTCTGCACCACCCCTCTGGTTCACACCGCACCCTGCACTCACCCACAGCAG CTCCATCCACTGGGAAAACTGGGGTTTGCATCACTCCACTGCACAGTGTTAGTGGGACCTGGGGGCAA GTCCCTTGACTTCTCTGAGCCTCAGTTTCCTTATGTGAAAGTTGCTGGAACCAAAATGGAGTCACTTA TGCCAAACTCTAATAAAATGGAGTCGGGGGGGCACATAGAAGCCCTCACACACACATGCCCGTAACAG GATTTATCACCAAGACACGCCTGCATGTAAGACCAGGACACACAGGGCGTATGGAAAAGCACGTCCTCAAA GACTGTAGTATTCCAGATGAGCTGCAGATGCTTACCTACCACGGCCGTCTCCACCAGAAAACCATCGC CAACTCCTGCGATCAGCTTGTGACTTACAAACCTTGTTTAAAAGCTGCTTACATGGACTTCTGTCCTT TAAAACGTTCCCCTTGGCTGTGGCCCTCTGTGTATGCCTGGGATCCTTCCAAGCACTCATAGCCCAGA TAGGAATCCTCTGCTCCTCCCAAATAAATTCATCTGTTC
	ORF Start: ATG at 617		ORF Stop: TGA at 1772

	SEQ ID NO: 20	385 aa	MW at 42794.8kD
NOV3a, CG118051-01 Protein Sequence	MKDEPRSTNLFMKLDSVFIWKEPFGLVLIIAPWNYPLNLTLVLLVGTLPAGNCVVLKPSEISQGTEKV LAEVLPQYLDQSCFAVVLGGPQETGQLLEHKLDYIFFTGSPRVGKIVMTAATKHLTPVTLELGGKNPC YVDDNCDPQTVANRVAWFCYFNAGQTCVAPDYVLCSPEMQERLLPALQSTITRFYGDDPQSSPNLGRI INQKQFQRLRALLGCGRVAIGGQSNESDRYIAPTVLVDVQETEPVMQEEIFGPILPIVNVQSVDEAIK FINRQEKPLALYAFSNSRQVVNQMLERTSSGSFGGNEGFTYISLLSVPFGGVGHSGMGRYHGKFTFDT FSHHRTCLLAPSGLEKLKEIRYPPYTDWNQQLLRWGMGSQSCTLL		

	SEQ ID NO: 21	1586 bp	
NOV3b, CG118051-02 DNA Sequence	CACGAGTTGGTTTGGGAGCTGCCAGTCTCCTGGGAGGATCGCAGTCAGCAGAGCAGGGCTGAGGCCT GGGGGTAGGAGCAGAGCCTGCGCATCTGGAGGCAGCATGTCCAAGAAAGGGAGTGGAGGTGCAGCGA AGGACCCAGGGGCAGAGCCCACGCTGGGGATGGACCCCTTCGAGGACACACTGCGGCGGCTGCGTGA GGCCTTCAACTGAGGGCGCACGCGGCCGGCCGAGTTCCGGGCTGCGCAGCTCCAGGGCCTGGGCCAC TTCCTTCAAGAAAACAAGCAGCTTCTGCGCGACGTGCTGGCCCAGGACCTGCATAAGCCAGCTTTCG AGGCAGACATATCTGAGCTCATCCTTTGCCAGAACGAGGTTGACTACGCTCTCAAGAACCTTCAGGC CTGGATGAAGGATGAACCACGGTCCACGAACCTGTTCATGAAGCTGGACTCGGTCTTCATCTGGAAG

	GAACCCTTTGGCCTGGTCCTCATCATCGCACCCTGGAACTACCCATTGAACCTGACCCTGGTGCTCC
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	TACTTCAATGCCGGCCAGACCTGCGTGGCCCCTGACTACGTCCTGTGCAGCCCCGAGATGCAGGAGA
	GGCTGCTGCCCGCCCTGCAGAGCACCATCACCCGTTTCTATGGCGACGACCCCCAGAGCTCCCCAAA
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	TGGAGAAATTAAAGGAGATCCGCTACCCACCCTATACCGACTGGAACCAGCAGCTGTTACGCTGGGG
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	AGCCTCAGGTTGCTGGAGCTGTCACATGACTGCATCCTGCCTGCC
ORF Start: ATG at 407			ORF Stop: TGA at 1436

	SEQ ID NO: 22	343 aa	MW at 38350.9kD
NOV3b, CG118051-02 Protein Sequence	MKDEPRSTNLFMKLDSVFIWKEPFGLVLIIAPWNYPLNLTLVLLVGTLPAGNCVVLKPSEISQGTEK VLAEVLPQYLDQSCFAVVLGGPQETGQLLEHKLDYIFFTGSPRVGKIVMTAATKHLTPVTLELGGKN PCYVDDNCDPQTVANRVAWFCYFNAGQTCVAPDYVLCSPEMQERLLPALQSTITRFYGDDPQSSPNL GRIINQKQFQRLRALLGCGRVAIGGQSNESDRYIAPTVLVDVQETEPVMQEEIFGPILPIVNVQSVD EAIKFINRQEKPLALHSGMGRYHGKFTFDTFSHHRTCLLAPSGLEKLKEIRYPPYTDWNQQLLRWGM GSQSCTLL		

	SEQ ID NO: 23	1791 bp	
NOV3c, CG118051-03 DNA Sequence	TTAAGGAGAATCTTAAAGTGAGGGCTGAGGGACTCTCCTGATCCAGAGCTGAGGACTCTCCTGATCCA GAGCTGAGGGCTCTCCTGATGACCCCTTCGAGGACACGCTGCGGCGGCTGCGTGAGGCCTTCAACTG AGGGCGCACGCGGCCGGCCGAGTTCCGGGCTGCGCAGCTCCAGGGCCTGGGCCCACTTCCTTCAAGAAA ACAAGCAGCTTCTGCGCACGTGCTGGCCCAGGACCTGCATAAGCCAGCTTTCGAGGCAGACATATCT GAGCTCATCCTTTGCCAGAACGAGGTTGACTACGCTCTCAAGAACCTTCAGGCCTGGATGAAGGATGA ACCACGGTCCACGAACCTGTTCATGAAGCTGGACTCGGTCTTCATCTGGAAGGAACCCTTTGGCCTGG TCCTCATCATCGCACCCTGGAACTACCCACTGAACCTGACCCTGGTGCTCCTGGTGGGCGCCCTCGCC GCAGGGAATTGCGTGGTGCTGAAGCCGTCAGAAATCAGCCAGGGCACAGAGAAGGTCCTGGCTGAGGT GCTGCCCCAGTACCTGGACCAGAGCTGCTTTGCCGTGGTGCTGGGCGGACCCCAGGAGACAGGGCAGC TGCTAGAGCACAAGTTGGACTACATCTTCTTCACAGGGGAGCCCTCGTGTGGGCAAGATTGTCATGACT GCTGCCACCAAGCACCTGACGCCTGTCACCCTGGAGCTGGGGGGCAAGAACCCCTGCTACGTGGACGA CAACTGCGACCCCCAGACCGTGGCCAACCGCGTGGCCTGGTTCTGCTACTTCAATGCCGGCCAGACCT GCGTGGCCCCTGACTACGTCCTGTGCAGCCCCGAGATGCAGGAGAGGCTGCTGCCCGCCCTGCAGAGC ACCATCACCCGTTTCTATGGCGACGACCCCCAGAGCTCCCCAAAACCTGGGCCGCATCATCAACCAGAA ACAGTTCCAGCGGCTGCGGGCATTGCTGGGCTGCGGCTGGGGGCCAGAGCAACGAGA GCGATCGCTACATCGCCCCCACGGTGCTGGTGGACGTGCAGGAGACGGAGCCTGTGATGCAGGAGGAG ATCTTCGGGCCCATCCTGCCCATCGTGAACGTGCAGAGCGTGGACGAGGCCATCAAGTTCATCAACCG GCAGGAGAAGCCCCTGGCCCTGTACGCCTTCTCCAACAGCAGCCAGGTTGTGAACCAGATGCTGGAGC GGGGGGAGTCGGCCACAGTGGGATGGGCCGGTACCACGGCAAGTTCACCTTCGACACCTTCTCCCACCA CCGCACCTGCCCGCTCGCCCCCTCCGGCCTGGAGAAATTAAAGGAGATCCGCTACCCACCCTATACCG ACTGGAACCAGCAGCTGTTACGCTGGGGCATGGGCTCCCAGAGCTGCACCCTCCTGTGAGCGTCCCAC CCGCCTCCAACGGGTCACACAGAGAAACCTGAGTCTAGCCATGAGGGGCTTATGCTCCCAACTCACAT TGTTCCTCCAGACCGCAGGCTCCCCCAGCCTCAGGGCTGCAAAGCAAGGTCTTGCTTCTATCTGGGGGGCGCTGCTCGAGAGAGGCCGAGAGGCCGC AGAACATGCCAGGTGTCCTCACTCACCCCACCCTCCCCAATTCCAGCCCTTTGCCCTCTCGGTCAGGG TTGACCAGGCCAAGGGCTAGCAT		
ORF Start: ATG at 330			ORF Stop: TGA at 1485

	SEQ ID NO: 24	385 aa	MW at 42653.5kD
NOV3c, CG118051-03 Protein Sequence	MKDEPRSTNLFMKLDSVFIWKEPFGLVLIIAPWNYPLNLTLVLLVGALAAGNCVVLKPSEISQGTEKV LAEVLPQYLDQSCFAVVLGGPQETGQLLEHKLDYIFFTGSPRVGKIVMTAATKHLTPVTLELGGKNPC YVDDNCDPQTVANRVAWFCYFNAGQTCVAPDYVLCSPEMQERLLPALQSTITRFYGDDPQSSPNLGRI INQKQFQRLRALLGCGRVAIGGQSNESDRYIAPTVLVDVQETEPVMQEEIFGPILPIVNVQSVDEAIK FINRQEKPLALYAFSNSSQVVNQMLERTSSGSFGGNEGFTYISLLSVPFGGVGHSGMGRYHGKFTFDT FSHHRTCPLAPSGLEKLKEIRYPPYTDWNQQLLRWGMGSQSCTLL		

Sequence comparison of the above protein sequences yields the following sequence relationships shown in Table 3B.

Table 3B. Comparison of NOV3a against NOV3b and NOV3c.		
Protein Sequence	NOV3a Residues/ Match Residues	Identities/ Similarities for the Matched Region
NOV3b	1..385 1..343	331/385 (85%) 331/385 (85%)
NOV3c	1..385 1..385	363/385 (94%) 363/385 (94%)

Further analysis of the NOV3a protein yielded the following properties shown in Table 3C.

Table 3C. Protein Sequence Properties NOV3a	
PSort analysis:	0.7900 probability located in plasma membrane; 0.3000 probability located in Golgi body; 0.2000 probability located in endoplasmic reticulum (membrane); 0.1743 probability located in microbody (peroxisome)
SignalP analysis:	Cleavage site between residues 54 and 55

A search of the NOV3a protein against the Geneseq database, a proprietary database that contains sequences published in patents and patent publication, yielded several homologous proteins shown in Table 3D.

Table 3D. Geneseq Results for NOV3a				
Geneseq Identifier	Protein/Organism/Length [Patent #, Date]	NOV3a Residues/	Identities/ Similarities for	Expect Value

		Match Residues	the Matched Region	
AAB58156	Lung cancer associated polypeptide sequence SEQ ID 494 - Homo sapiens, 430 aa. [WO200055180-A2, 21-SEP-2000]	1..353 62..414	325/353 (92%) 337/353 (95%)	0.0
ABB66868	Drosophila melanogaster polypeptide SEQ ID NO 27396 - Drosophila melanogaster, 561 aa. . [WO200171042-A2, 27-SEP-2001]	14..309 95..390	158/296 (53%) 212/296 (71%)	3e-94
ABB65492	Drosophila melanogaster polypeptide SEQ ID NO 23268 - Drosophila melanogaster, 561 aa. [WO200171042-A2, 27-SEP-2001]	14..309 95..390	158/296 (53%) 212/296 (71%)	3e-94
ABP39856	Staphylococcus epidermidis ORF amino acid sequence SEQ ID NO:4701 - Staphylococcus epidermidis, 464 aa. [US6380370-B1, 30-APR-2002]	2..365 88..451	157/366 (42%) 235/366 (63%)	1e-85
AAG82730	S. epidermidis open reading frame protein sequence SEQ ID NO:2554 - Staphylococcus epidermidis, 459 aa. [WO200134809-A2, 17-MAY-2001]	2..365 83..446	157/366 (42%) 235/366 (63%)	1e-85

In a BLAST search of public sequence datbases, the NOV3a protein was found to have homology to the proteins shown in the BLASTP data in Table 3E.

5

Table 3E. Public BLASTP Results for NOV3a				
Protein Accession Number	Protein/Organism/Length	NOV3a Residues/ Match Residues	Identities/ Similarities for the Matched Portion	Expect Value
P48448	Aldehyde dehydrogenase 8 (EC 1.2.1.5) - Homo sapiens (Human), 385 aa.	1..385 1..385	385/385 (100%) 385/385 (100%)	0.0

BAC03897	CDNA FLJ35145 fis, clone PLACE6009853, highly similar to ALDEHYDE DEHYDROGENASE 8 (EC 1.2.1.5) - Homo sapiens (Human), 385 aa.	1..385 1..385	380/385 (98%) 381/385 (98%)	0.0
P43353	Aldehyde dehydrogenase 7 (EC 1.2.1.5) - Homo sapiens (Human), 468 aa.	1..385 82..468	321/387 (82%) 345/387 (88%)	0.0
AAH33099	Similar to aldehyde dehydrogenase 3 family, member B1 - Homo sapiens (Human), 431 aa.	13..385 57..431	315/375 (84%) 339/375 (90%)	0.0
Q8VHW0	Aldehyde dehydrogenase ALDH3B1 (EC 1.2.1.3) - Mus musculus (Mouse), 449 aa (fragment).	1..385 63..449	295/387 (76%) 336/387 (86%)	e-174

PFam analysis predicts that the NOV3a protein contains the domains shown in the Table 3F.

5

Table 3F. Domain Analysis of NOV3a			
Pfam Domain	**NOV3a Match Region**	**Identities/ Similarities for the Matched Region**	**Expect Value**
aldedh	1..351	129/492 (26%) 299/492 (61%)	1.1e-103

Example 4.

10   The NOV4 clone was analyzed, and the nucleotide and encoded polypeptide sequences are shown in Table 4A.

Table 4A. NOV4 Sequence Analysis			
	SEQ ID NO: 25	1636 bp	
NOV4a, CG120277-01 DNA Sequence	CCAGGAGCCCCAGTTACCGGGAGAGGCTGTGTCAAAGGCGCCATGAGCAAGATCAGCGAGGCCGTGAA GCGCGCCCGCGCCGCCTTCAGCTCGGGCAGGACCCGTCCGCTGCAGTTCCGATTCCAGCAGCTGGAGG CGCTGCAGCGCCTGATCCAGGAGCAGGAGCAGGAGCTGGTGGGCGCGCTGGCCGCAGACCTGCACAAG AATGAATGGAACGCCTACTATGAGGAGGTGGTGTACGTCCTAGAGGAGATCGAGTACATGATCCAGAA GCTCCCTGAGTGGGCCGCGGATGAGCCCGTGGAGAAGACGCCCCAGACTCAGCAGGACGAGCTCTACA TCCACTCGGAGCCACTGGGCGTGGTCCTCGTCATTGGCACCTGGAACTACCCCTTCAACCTCACCATC CAGCCCATGGTGGGCGCCATCGCTGCAGGGAACGCAGTGGTCCTCAAGCCCTCGGAGCTGAGTGAGAA CATGGCGAGCCTGCTGGCTACCATCATCCCCCAGTACCTGGACAAGGATCTGTACCCAGTAATCAATG		

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CG120277-01
Protein
Sequence</td><td colspan="3">MSKISEAVKRARAAFSSGRTRPLQFRFQQLEALQRLIQEQEQELVGALAADLHKNEWNAYYEEVVYVL
EEIEYMIQKLPEWAADEPVEKTPQTQQDELYIHSEPLGVVLVIGTWNYPFNLTIQPMVGAIAAGNAVV
LKPSELSENMASLLATIIPQYLDKDLYPVINGGVPETTELLKERFDHILYTGSTGVGKIIMTAAAKHL
TPVTLELGGKSPCYVDKNCDLDVACRRIAWGKFMNSGQTCVAPDYILCDPSIQNQIVEKLKKSLKEFY
GEDAKKSRDYGRIISARHFQRVMGLIEGQKVAYGGTGDAATRYIAPTILTDVDPQSPVMQEEIFGPVL
PIVCVRSLEEAIQFINQREKPLALYMFSSNDKVIKKMIAETSSGGVAANDVIVHITLHSLPFGGVGNS
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<tr><td>NOV4b,
CG120277-02
DNA Sequence</td><td colspan="3">GAGCCCCAGTTACCGGGAGAGGCTGTGTCAAAGGCGCC**ATG**AGCAAGATCAGCGAGGCCGTGAAGCG
CGCCCGCGCCGCCTTCAGCTCGGGCAGGACCCGTCCGCTGCAGTTCCGGATCCAGCAGCTGGAGGCG
CTGCAGCGCCTGATCCAGGAGCAGGAGCAGGAGCTGGTGGGCGCGCTGGCCGCAGACCTGCACAAGA
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TGGGGGAAATTCATGAACAGTGGCCAGACCTGCTGTGGCCCCAGACTACATCCTCTGTGACCCCTCGA
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CAGAAGGTGGCTTATGGGGGCACCGGGGATGCCGCCACTCGCTACATAGCCCCCACCATCCTCACGG
ACGTGGACCCCCAGTCCCCGGTGATGCAAGAGGAGATCTTCGGGCCTGTGCTGCCCATCGTGTGCGT
GCGCAGCCTGGAGGAGGCCATCCAGTTCATCAACCAGCGTGAGAAGCCCCTGGCCCTCTACATGTTC
TCCAGCAACGACAAGGTGATTAAGAAGATGATTGCAGAGACATCCAGTGGTGGGGTGGCGGCCAACG
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GCCTCTGATGAATGATGAAGGCCTGAAGGTCAGATACCCCCCGAGCCCGGCCAAGATGACCCAGCAC
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	SEQ ID NO: 28	434 aa	MW at 48169.0kD
NOV4b, CG120277-02 Protein Sequence	MSKISEAVKRARAAFSSGRTRPLQFRIQQLEALQRLIQEQEQELVGALAADLHKNEWNAYYEEVVYV LEEIEYMIQKLPEWAADEPVEKTPQTQQDELYIHSEPLGVVLVIGTWNYPFNLTIQPMVGAIAAGNA VVLKPSELSENMASLLATIIPQYLDKDLYPVINGGVPETTELLKERFDHILYTGSTGVGKIIMTAAA KHLTPVTLELGGKSPCYVDKNCDLDVACRRIAWGKFMNSGQTCVAPDYILCDPSIQNQIVEKLKKSL KEFYGEDAKKSRDYGRIISARHFQRVMGLIEGQKVAYGGTGDAATRYIAPTILTDVDPQSPVMQEEI FGPVLPIVCVRSLEEAIQFINQREKPLALYMFSSNDKVIKKMIAETSSGGVAANDVIVHITLHSLPF GGVGNSGMVRPLMNDEGLKVRYPPSPAKMTQH		

Sequence comparison of the above protein sequences yields the following sequence relationships shown in Table 4B.

Table 4B. Comparison of NOV4a against NOV4b.		
Protein Sequence	NOV4a Residues/ Match Residues	Identities/ Similarities for the Matched Region
NOV4b	1..453 1..434	401/453 (88%) 401/453 (88%)

Further analysis of the NOV4a protein yielded the following properties shown in Table 4C.

Table 4C. Protein Sequence Properties NOV4a	
PSort analysis:	0.7636 probability located in mitochondrial matrix space; 0.4422 probability located in mitochondrial inner membrane; 0.4422 probability located in mitochondrial intermembrane space; 0.4422 probability located in mitochondrial outer membrane
SignalP analysis:	No Known Signal Sequence Predicted

A search of the NOV4a protein against the Geneseq database, a proprietary database that contains sequences published in patents and patent publication, yielded several homologous proteins shown in Table 4D.

Table 4D. Geneseq Results for NOV4a				
Geneseq Identifier	Protein/Organism/Length [Patent #, Date]	NOV4a Residues/ Match	Identities/ Similarities for the Matched	Expect Value

		Residues	Region	
AAB58156	Lung cancer associated polypeptide sequence SEQ ID 494 - Homo sapiens, 430 aa. [WO200055180-A2, 21-SEP-2000]	48..431 28..411	208/384 (54%) 277/384 (71%)	e-124
ABB66868	Drosophila melanogaster polypeptide SEQ ID NO 27396 - Drosophila melanogaster, 561 aa. [WO200171042-A2, 27-SEP-2001]	1..394 1..394	199/394 (50%) 270/394 (68%)	e-115
ABB65492	Drosophila melanogaster polypeptide SEQ ID NO 23268 - Drosophila melanogaster, 561 aa. [WO200171042-A2, 27-SEP-2001]	1..394 1..394	199/394 (50%) 270/394 (68%)	e-115
AAG21988	Arabidopsis thaliana protein fragment SEQ ID NO: 24747 - Arabidopsis thaliana, 484 aa. [EP1033405-A2, 06-SEP-2000]	2..445 10..456	210/449 (46%) 288/449 (63%)	e-112
AAG11789	Arabidopsis thaliana protein fragment SEQ ID NO: 10644 - Arabidopsis thaliana, 484 aa. [EP1033405-A2, 06-SEP-2000]	2..445 10..456	210/449 (46%) 288/449 (63%)	e-112

In a BLAST search of public sequence datbases, the NOV4a protein was found to have homology to the proteins shown in the BLASTP data in Table 4E.

5

Table 4E. Public BLASTP Results for NOV4a				
Protein Accession Number	Protein/Organism/Length	NOV4a Residues/ Match Residues	Identities/ Similarities for the Matched Portion	Expect Value
P30838	Aldehyde dehydrogenase, dimeric NADP-preferring (EC 1.2.1.5) (ALDH class 3) (ALDHIII) - Homo sapiens (Human), 453 aa.	1..453 1..453	453/453 (100%) 453/453 (100%)	0.0

Q9BT37	Aldehyde dehydrogenase 3 (Aldehyde dehydrogenase 3 family, member A1) - Homo sapiens (Human), 453 aa.	1..453 1..453	452/453 (99%) 452/453 (99%)	0.0
A42584	aldehyde dehydrogenase (NAD(P)+) (EC 1.2.1.5) 3 - human, 453 aa.	1..453 1..453	450/453 (99%) 451/453 (99%)	0.0
A30149	aldehyde dehydrogenase (NADP+) (EC 1.2.1.4) 3, tumor-associated [similarity] - rat, 453 aa.	1..453 1..453	370/453 (81%) 415/453 (90%)	0.0
P11883	Aldehyde dehydrogenase, dimeric NADP-preferring (EC 1.2.1.5) (ALDH class 3) (Tumor-associated aldehyde dehydrogenase) (HTC-ALDH) - Rattus norvegicus (Rat), 452 aa.	2..453 1..452	369/452 (81%) 414/452 (90%)	0.0

PFam analysis predicts that the NOV4a protein contains the domains shown in the Table 4F.

5

Table 4F. Domain Analysis of NOV4a			
Pfam Domain	**NOV4a Match Region**	**Identities/ Similarities for the Matched Region**	**Expect Value**
aldedh	1..432	182/492 (37%) 401/492 (82%)	7.4e-206

Example 5.

10   The NOV5 clone was analyzed, and the nucleotide and encoded polypeptide sequences are shown in Table 5A.

Table 5A. NOV5 Sequence Analysis			
	SEQ ID NO: 29	2316 bp	
NOV5a, CG140468-01 DNA Sequence	GCCACGAAGGCCACAGACGCCTTCCCCCTTGGACTCTCATTCCCTTTTTCCACGGAGCCCCGCGCTTTC GTGAGCCCCCTCGAGGAACCTGGTCTCCGCATCCAGTTACCACCTCCTGCCTCAGAGGCCATCTGAGC CCTTCGCACCTCGCCCCTCAGTCCCCCCTTGCCCCCCCCGCGGAGATCGCCTCGCTCCCTCCCGCCCCC CCATCATCCCTTCCCTCGCAGTTCCCCTGTCCTGAGGGGAGCCCCGCCACGGCAGCGACAGCGGGCAG GAGGGAGAAAGTGAAGGTTGGGCGACACTTGGCCTCACTCCCGGCTAGGCGCACCCACGGGGAGGAGA GGAGGAGCCGAGAGAGCTGAGCAGCGCGGAAGTAGCTGCTGCTGGTGGTGACAATGTCAAATAACGGC CTAGACATTCAAGACAAACCCCCAGCCCCTCCGATGAGAAATACCAGCACTATGATTGGAGTCGGCAG		

	CAAAGATGCTGGAACCCTAAACCATGGTTCTAAACCTCTGCCTCCAAACCCAGAGGAGAAGAAAAAGA
	AGGACCGATTTTACCGATCCATTTTACCTGGAGATAAAACAAATAAAAAGAAAGAGAAAGAGCGGCCA
	GAGATTTCTCTCCCTTCAGATTTTGAACACACAATTCATGTCGGTTTTGATGCTGTCACAGGGGAGTT
	TACGGGAATGCCAGAGCAGTGGGCCGCTTGCTTCAGACATCAAATATCACTAAGTCGGAGCAGAAGA
	AAAACCCGCAGGCTGTTCTGGATGTGTTGGAGTTTTACAACTCGAAGAAGACATCCAACAGCCAGAAA
	TACATGAGCTTTACAGATAAGTCAGCTGAGGATTACAATTCTTCTAATGCCTTGAATGTGAAGGCTGT
	GTCTGAGACTCCTGCAGTGCCACCAGTTTCAGAAGATGAGGATGATGATGATGATGCTACCCCAC
	CACCAGTGATTGCTCCACGCCCAGAGCACACAAAATCTGTATACACACGGTCTGTGATTGAACCACTT
	CCTGTCACTCCAACTCGGGACGTGGCTACATCTCCCATTTCACCTACTGAAAATAACACCACTCCACC
	AGATGCTTTGACCCGGAATACTGAGAAGCAGAAGAAGAAGCCTAAAATGTCTGATGAGGAGATCTTGG
	AGAAATTACGAAGCATAGTGAGTGTGGGCGATCCTAAGAAGAAATATACACGGTTTGAGAAGATTGGA
	CAAGGTGCTTCAGGCACCGTGTACACAGCAATGGATGTGGCCACAGGACAGGAGGTGGCCATTAAGCA
	GATGAATCTTCAGCAGCAGCCCAAGAAAGAGCTGATTATTAATGAGATCCTGGTCATGAGGGAAAACA
	AGAACCCAAACATTGTGAATTACTTGGACAGTTACCTCGTGGGAGATGAGCTGTGGGTTGTTATGGAA
	TACTTGGCTGGAGGCTCCTTGACAGATGTGGTGACAGAAACTTGCATGGATGAAGGCAAATTGCAGC
	TGTGTGCCGTGAGTGTCTGCAGGCTCTGGAGTTCTTGCATTCGAACCAGGTCATTCACAGAGACATCA
	AGAGTGACAATATTCTGTTGGGAATGGATGGCTCTGTCAAGCTAACTGACTTTGGATTCTGTGCACAG
	ATAACCCCAGAGCAGAGCAAACGGAGCACCATGGTAGGAACCCCATACTGGATGGCACCAGAGGTTGT
	GACACGAAAGGCCTATGGGCCCAAGGTTGACATCTGGTCCCTGGGCATCATGGCCATCGAAATGATTG
	AAGGGGAGCCTCCATACCTCAATGAAAACCCTCTGAGAGCCTTGTACCTCATTGCCACCAATGGGACC
	CCAGAACTTCAGAACCCAGAGAAGCTGTCAGCTATCTTCCGGGACTTTCTGAACCGCTGTCTCGATAT
	GGATGTGGAGAAGAGAGGTTCAGCTAAAGAGCTGCTACAGCATCAATTCCTGAAGATTGCCAAGCCCT
	TCTCCAGCCTCACTCCACTGATTGCTGCAGCTAAGGAGGCAACAAAGAACAATCAC**TAA**AACCACACT
	CACCCCAGCCTCATTGTGCCAAGCTCTGTGAGATAAATGCACATTTCAGAAATTCCAACTCCTGATGC
	CCTCTTCTCCTTGCCTTGCTTCTCCCATTTCCTGATCTAGCACTCCTCAAGACTTTGATCCTTGGAAA
	CCGTGTGTCCAGCATTGAAGAGAACTGCAACTGAATGACTAATCAGGATGATGGCCATTTCTAAATAAG
	GAATTTCCTCCCAATTCATGGATATGAGGGTGGTTTATGATTAAGGGTTTATATAAATAAATGTTTCT
	AGTC
ORF Start: ATG at 394		**ORF Stop: TAA at 2029**

	SEQ ID NO: 30	545 aa	MW at 60660.3kD
NOV5a, CG140468-01 Protein Sequence	MSNNGLDIQDKPPAPPMRNTSTMIGVGSKDAGTLNHGSKPLPPNPEEKKKKDRFYRSILPGDKTNKKK EKERPEISLPSDFEHTIHVGFDAVTGEFTGMPEQWARLLQTSNITKSEQKKNPQAVLDVLEFYNSKKT SNSQKYMSFTDKSAEDYNSSNALNVKAVSETPAVPPVSEDEDDDDDDATPPPVIAPRPEHTKSVYTRS VIEPLPVTPTRDVATSPISPTENNTTPPDALTRNTEKQKKKPKMSDEEILEKLRSIVSVGDPKKKYTR FEKIGQGASGTVYTAMDVATGQEVAIKQMNLQQQPKKELIINEILVMRENKNPNIVNYLDSYLVGDEL WVVMEYLAGGSLTDVVTETCMDEGQIAAVCRECLQALEFLHSNQVIHRDIKSDNILLGMDGSVKLTDF GFCAQITPEQSKRSTMVGTPYWMAPEVVTRKAYGPKVDIWSLGIMAIEMIEGEPPYLNENPLRALYLI ATNGTPELQNPEKLSAIFRDFLNRCLDMDVEKRGSAKELLQHQFLKIAKPLSSLTPLIAAAKEATKNN H		

	SEQ ID NO: 31	957 bp	
NOV5b, CG140468-02 DNA Sequence	GACAATGTCAAATAACGGCCTAGCATTCAAGACAAACCCCCAGCCCCTCCGATGAGAAATACCAGC ACTATGATTGGAGCCGGCAGCAAAGATGCTGGAACCCTAAACCATGGTTCTAAACCTCTGCCTCCAA ACCCAGAGGAGAAGAAAAAGAAGGACCGATTTTACCGATCCATTTTACCTGGAGATAAAACAAATAA AAAGAAAGAGAAAGAGCGGCCAGAGATTTCTCTCCCTTCAGATTTTGAACACACAATTCATGTCGGT TTTGATGCTGTCACAGGGGAGTTTACGGGAATGCCAGAGCAGTGGGCCCGCTTGCTTCAGACATCAA GAAGAAGACATCCAACAGCCAGAAATACATGAGCTTTACAGATAAGTCAGCTGAGGATTACAATTCT TCTAATGCCTTGAATGTGAAGGCTGTGTCTGAGACTCCTGCAGTGCCACCAGTTTCAGAAGATGAGG ATGATGATGATGATGCTACCCCACCACCAGTGATTGCTCCACGCCCAGAGCACACAAAATCTGT ATACACACGGTCTGTGATTGAACCACTTCCTGTCACTCCAACTCGGGACGTGGCTACATCTCCCATT TCACCTACTGAAAATAACACCACTCCACCAGATGCTTTGACCCGGAATACTGAGAAGCAGAAGAAGA AGCCTAAAATGTCTGATGAGGAGATCTTGGAGAAATTACGAAGCATAGTGAGTGTGGGCGATCCTAA GAAGAAATATACACGGTTTGAGAAGATTGCCAAGCCCCTCTCCAGCCTCACTCCACTGATTGCTGCA GCTAAGGAGGCAACAAAGAACAATCAC**TAA**AACCACACTCACCCCAGCCTCATTGTGCCAAGCCTTC TGTGAGATAAATGCACATT		
	ORF Start: ATG at 5		**ORF Stop: TAA at 899**

	SEQ ID NO: 32	298 aa	MW at 32989.7kD
NOV5b, CG140468-02 Protein Sequence	MSNNGLDIQDKPPAPPMRNTSTMIGAGSKDAGTLNHGSKPLPPNPEEKKKKDRFYRSILPGDKTNKK KEKERPEISLPSDFEHTIHVGFDAVTGEFTGMPEQWARLLQTSNITKSEQKKNPQAVLDVLEFYNSK KTSNSQKYMSFTDKSAEDYNSSNALNVKAVSETPAVPPVSEDEDDDDDDATPPPVIAPRPEHTKSVY TRSVIEPLPVTPTRDVATSPISPTENNTTPPDALTRNTEKQKKPKMSDEEILEKLRSIVSVGDPKK KYTRFEKIAKPLSSLTPLIAAAKEATKNNH		

Sequence comparison of the above protein sequences yields the following sequence relationships shown in Table 5B.

Table 5B. Comparison of NOV5a against NOV5b.		
Protein Sequence	**NOV5a Residues/ Match Residues**	**Identities/ Similarities for the Matched Region**
NOV5b	1..281 1..281	238/281 (84%) 239/281 (84%)

Further analysis of the NOV5a protein yielded the following properties shown in Table 5C.

Table 5C. Protein Sequence Properties NOV5a	
PSort analysis:	0.7000 probability located in nucleus; 0.3000 probability located in microbody (peroxisome); 0.1000 probability located in mitochondrial matrix space; 0.1000 probability located in lysosome (lumen)
SignalP analysis:	No Known Signal Sequence Predicted

A search of the NOV5a protein against the Geneseq database, a proprietary database that contains sequences published in patents and patent publication, yielded several homologous proteins shown in Table 5D.

Table 5D. Geneseq Results for NOV5a				
Geneseq Identifier	**Protein/Organism/Length [Patent #, Date]**	**NOV5a Residues/ Match**	**Identities/ Similarities for the Matched**	**Expect Value**

		Residues	Region	
AAB03968	p-21 activated protein kinase (PAK1) - Homo sapiens, 545 aa. [WO200060062-A2, 12-OCT-2000]	1..545 1..545	544/545 (99%) 545/545 (99%)	0.0
AAY55958	Human STE20-related protein kinase PAK1_h - Homo sapiens, 545 aa. [WO9953036-A2, 21-OCT-1999]	1..545 1..545	541/545 (99%) 542/545 (99%)	0.0
ABG30251	Novel human diagnostic protein #30242 - Homo sapiens, 587 aa. [WO200175067-A2, 11-OCT-2001]	1..542 7..557	474/556 (85%) 500/556 (89%)	0.0
AAW72757	Human doublin - Homo sapiens, 544 aa. [WO9840495-A1, 17-SEP-1998]	3..544 2..542	444/552 (80%) 489/552 (88%)	0.0
ABB57290	Mouse ischaemic condition related protein sequence SEQ ID NO:817 - Mus musculus, 544 aa. [WO200188188-A2, 22-NOV-2001]	3..544 2..542	441/552 (79%) 483/552 (86%)	0.0

In a BLAST search of public sequence datbases, the NOV5a protein was found to have homology to the proteins shown in the BLASTP data in Table 5E.

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Table 5E. Public BLASTP Results for NOV5a				
Protein Accession Number	Protein/Organism/Length	NOV5a Residues/ Match Residues	Identities/ Similarities for the Matched Portion	Expect Value
Q13153	Serine/threonine-protein kinase PAK 1 (EC 2.7.1.-) (p21-activated kinase 1) (PAK-1) (P65-PAK) (Alpha-PAK) - Homo sapiens (Human), 545 aa.	1..545 1..545	545/545 (100%) 545/545 (100%)	0.0

P35465	Serine/threonine-protein kinase PAK 1 (EC 2.7.1.-) (p21-activated kinase 1) (PAK-1) (P68-PAK) (Alpha-PAK) (Protein kinase MUK2) - Rattus norvegicus (Rat), 544 aa.	1..545 1..544	537/545 (98%) 539/545 (98%)	0.0
S40482	serine/threonine-specific protein kinase (EC 2.7.1.-) - rat, 544 aa.	1..545 1..544	534/545 (97%) 537/545 (97%)	0.0
O88643	Serine/threonine-protein kinase PAK 1 (EC 2.7.1.-) (p21-activated kinase 1) (PAK-1) (P65-PAK) (Alpha-PAK) (CDC42/RAC effector kinase PAK-A) - Mus musculus (Mouse), 545 aa.	1..545 1..545	530/545 (97%) 537/545 (98%)	0.0
O75561	P21 activated kinase 1B - Homo sapiens (Human), 553 aa.	1..522 1..522	517/522 (99%) 520/522 (99%)	0.0

PFam analysis predicts that the NOV5a protein contains the domains shown in the Table 5F.

Table 5F. Domain Analysis of NOV5a			
Pfam Domain	NOV5a Match Region	Identities/ Similarities for the Matched Region	Expect Value
PBD	75..135	37/64 (58%) 59/64 (92%)	3.4e-34
pkinase	270..521	94/291 (32%) 208/291 (71%)	5.7e-90

Example 6.

The NOV6 clone was analyzed, and the nucleotide and encoded polypeptide sequences are shown in Table 6A.

Table 6A. NOV6 Sequence Analysis			
	SEQ ID NO: 33	3255 bp	

| NOV6a, CG142182-01 DNA Sequence | GACAGCTTTGGGTGGACCAGTAATGAGGAAATGAGGCAACATGATGTGCAGGAACTGAATCGAATCCT CTTCAGCGCTTTGGAAACTTCTTTAGTTGGGACCTCCGGTCATGACCTCATCTATCGTCTGTACCATG GAACCATTGTTAACCAGATTGTTTGTAAAGAATGTAAGAACGTTAGCGAGAGGCAGGAAGACTTCTTA GATCTAACAGTAGCAGTCAAAAATGTATCCGGTTTGGAAGATGCTCTCTGGAACATGTATGTAGAAGA GGAAGTTTTTGATTGTGACAACTTGTACCACTGTGGAACTTGTGACAGGCTGGTTAAAGCAGCAAAGT CGGCCAAATTACGTAAGCTGCCTCCTTTTCTTACTGTTTCATTACTAAGATTTAATTTTGATTTTGTG AAATGCGAACGCTACAAGGAAACTAGCTGCTGTTATACATTCCCTCTCCGGATTAATCTCAAGCCCTTTTG TGAACAGAGTGAATTGGATGACTTAGAATATATATATGACCTCTTCTCAGTTATTATACACAAAGGTG GCTGCTACGGAGGCCATTACCATGTATATATTAAAGATGTTGATCATTTGGGAAACTGGCAGTTTCAA GAGGAAAAAAGTAAACCAGATGTGAATCTGAAAGATCTCCAGAGTGAAGAAGAGATTGATCATCCACT GATGATTCTAAAAGCAATCTTATTAGAGGAGGAGAATAATCTAATTCCTGTTGATCAGCTGGGCCAGA AACTTTTGAAAAAGATAGGAATATCTTGGAACAAGAAGTACAGAAAACAGCCATGGGACCATTGCGGAAG TTCTTACAGCTCCATTCTCAGATATTTCTACTCAGTTCAGATGAAAGTACAGTTCGTCTCTTGAAGAA TAGTTCTCTCCAGGCTGAGTCTGATTTCCAAAGGAATGACCAGCAAATTTTCAAGATGCTTCCTCCAG AATCCCCAGGTTTAAACAATAGCATCTCCTGTCCCCACTGGTTTGATATAAATGATTCTAAAGTCCAG CCAATCAGGGAAAAGGGATATTGAACAGCAATTTCAGGGTAAAGAAAGTGCCTACATGTTGTTTTATCG GAAATCCCAGTTGCAGAGACCCCCTGAAGCTCGAGCTAATCCAAGATATGGGGTTCCATGTCATTTAC TGAATGAAATGGATGCAGCTAACATTGAACTGCAAACCAAAAGGGCAGAATGTGATTCTGCAAACAAT ACTTTTGAATTGCATCTTCACCTGGGCCCTCAGTATCATTTCTTCAATGGGGCTCTGCACCCAGTAGT CTCTCAAACAGAAAGCGTGTGGGATTTGACCTTTGATAAAAGAAAAACTTTAGGAGATCTCCGGCAGT CAATATTTCAGCTGTTAGAATTTTTGGGAAGGAGACATGGTTCTTAGTGTTGCAAAGCTTGTACCAGCA GGACTTCACATTTACCAGTCACTTGGCGGGGATGAACTGACACTGTGTGAAACTGAAATTGCTGATGG GGAAGACATCTTTGTGTGGAATGGGGTGGAGGTTGGTGGAGTCCACATTCAAACTGGTATTGACTGCG AACCTCTACTTTTTAAATGTTCTTCATCTAGACACAAGCAGTGATGGAGAAAAGTGTTGTCAGGTGATA GAATCTCCACATGTCTTTCCAGCTAATGCAGAAGTGGGCACTGTCCTCAACAGCCTTAGCAATCCCAGC AGGTGTCATCTTCATCAACAGTGCTGGATGTCCAGGTGGGGAGGGTTGGACGGCCATCCCCAAGGAAG ACATGAGGAAGACGTTCAGGGAGCAAGGGCTCAGAAATGGAAGCTCAATTTTAATTCAGGATTCTCAT GATGATAACAGCTTGTTGACCAAGGAAGAGAAATGGGTCACTAGTATGAATGAGATTGACTGGCTCCA CGTTAAAAATTTATGCCAGTTAGAATCTGAAGAGAAGCAAGTTAAAATATCAGCAACTGTTAACACAA TGGTGTTTGATATTCGAATTAAAGCCATAAAGGAATTAAAATTAATGAAGGAACTAGCTGACAACAGC AGCAGAATTGAAGATGGGAAGTTCATTGGGACTGTGTCTTGGAAAAAGCACCAAGTTCGTCTCAGTTGT TCCTGTTTTTTGCAATGGGGAGTGACGTTCAACCTGGGACAGAAATGGAACTGACCGGCAGCCACTCA ATATCTGTGAGAGATTGTTTAAAGTTAATGCTGAAGAAATCTGGCCTACAAGACTCCTTTATAGGAGA TGCCTGGCATTTACGAAAAATGGATTGGTGCTATGAAGCTGGAGAGCCTTTATGTGAAGAAGATGCAA CACTGAAAGAACTTCTGATATGTTCTGGAGATACTTTGCTTTTAATTGAAGGACAACTTCCTCCTCTG GGTTTCCTGAAGGTGCCCATCTGGTGGTACCAGCTTCAGGGTCCCTCAGGACACTGGGAGAGTCATCA GGACCAGACCAACTGTACTTCGTCTTGGGGCAGAGTTTGGAGAGCCACTTCCAGCCAAGGTGCTTCTG GGAACGAGCCTGCGCAAGTTTCTCTCCTCTACTTGGGAGACATAGAGATCTCAGAAGATGCCACGCTG GCGGAGCTGAAGTCTCAGGCCATGACCTTGCCTCCTTTCCTGGAGTTCGGTGTCCCGTCCCAGCCCA CCTCAGAGCCTGGACGGTGGAGGAGGAAGCGCCCAGGCAGGCTTTTACGAACTGACCGGCAGCCACTCA GGGAATATAAACTAGGACGGAGAATTGAGATCTGCTTAGAGCCCCCTTCAGAAAAGGCGAAAACTTGGGC CCCCAGGACGTGCTGCTGAGGACACAGGTGCGCATCCCTGGTGAGAGGACCTATGCCCCTGCCCTGGA CCTGGTGTGGAACGCGGCCCAGGGTGGGACTGCCGGCTCCCTGAGGCAGAGAGTTGCCGATTTCTATT GTCTTCCCGTGGAGAAGATTGAAATTGCCAAATACTTTCCCGAAAAGTTCGAGTGGCTTCCGATATCT AGCTGGAACCAACAAATAACCAAGAGGAAAAAAAAAAAAAAAAACAAGATTATTTGCAAGGGGCACCGTA TTACTTGAAAGACGGAGATACTATTGGTGTTAAGGTAAGTTGTTTAACAGCAAATTTACCACTTTTGAG AAGACACGAGGGTCACATGATTTTATAGAGACGTTTTATTGAATCTTCAAGACACAGAT |

ORF Start: ATG at 31			ORF Stop: TGA at 3193

	SEQ ID NO: 34	1054 aa	MW at 119613.5kD
NOV6a, CG142182-01 Protein Sequence	MRQHDVQELNRILFSALETSLVGTSGHDLIYRLYHGTIVNQIVCKECKNVSERQEDFLDLTVAVKNVS GLEDALWNMYVEEEVFDCDNLYHCGTCDRLVKAAKSAKLRKLPPFLTVSLLRFNFDFVKCERYKETSC YTFPLRINLKPFCEQSELDDLEYIYDLFSVIIHKGGCYGGHYHVYIKDVDHLGNWQFQEEKSKPDVNL KDLQSEEEIDHPLMILKAILLEEENNLIPVDQLGQKLLKKIGISWNKKYRKQHGPLRKFLQLHSQIFL LSSDESTVRLLKNSSLQAESDFQRNDQQIFKMLPPESPGLNNSISCPHWFDINDSKVQPIREKDIEQQ FQGKESAYMLFYRKSQLQRPPEARANPRYGVPCHLLNEMDAANIELQTKRAECDSANNTPELHLHLGP QYHFFNGALHPVVSQTESVWDLTFDKRKTLGDLRQSIFQLLEFWEGDMVLSVAKLVPAGLHIYQSLGG DELTLCETEIADGEDIFVWNGVEVGGVHIQTGIDCEPLLLNVLHLDTSSDGEKCCQVIESPHVFPANA EVGTVLTALAIPAGVIFINSAGCPGGEGWTAIPKEDMRKTFREQGLRNGSSILIQDSHDDNSLLTKEE KWVTSMNEIDWLHVKNLCQLESEEKQVKISATVNTMVFDIRIKAIKELKLMKELADNSCLRPIDRNGK LLCPVPDSYTLKEAELKMGSSLGLCLGKAPSSSQLFLFFAMGSDVQPGTEMEIVVEETISVRDCLKLM LKKSGLQDSFIGDAWHLRKMDWCYEAGEPLCEEDATLKELLICSGDTLLLIEGQLPPLGFLKVPIWWY QLQGPSGHWESHQDQTNCTSSWGRVWRATSSQGASGNEPAQVSLLYLGDIEISEDATLAELKSQAMTL PPFLEFGVPSPAHLRAWTVERKRPGRLLRTDRQPLREYKLGRRIEICLEPLQKGENLGPQDVLLRTQV RIPGERTYAPALDLVWNAAQGGTAGSLRQRVADFYCLPVEKIEIAKYFPEKFEWLPISSWNQQITKRK KKKKQDYLQGAPYYLKDGDTIGVKVSCLTANLPL

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Further analysis of the NOV6a protein yielded the following properties shown in Table 6B.

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Table 6B. Protein Sequence Properties NOV6a	
PSort analysis:	0.7000 probability located in plasma membrane; 0.3500 probability located in nucleus; 0.3000 probability located in microbody (peroxisome); 0.2000 probability located in endoplasmic reticulum (membrane)
SignalP analysis:	No Known Signal Sequence Predicted

A search of the NOV6a protein against the Geneseq database, a proprietary database that contains sequences published in patents and patent publication, yielded

10   several homologous proteins shown in Table 6C.

Table 6C. Geneseq Results for NOV6a				
Geneseq Identifier	Protein/Organism/Length [Patent #, Date]	NOV6a Residues/ Match Residues	Identities/ Similarities for the Matched Region	Expect Value
AAE14346	Human protease PRTS-11 protein - Homo sapiens, 1108 aa. [WO200183775-A2, 08-NOV-2001]	1..1044 1..1040	1037/1044 (99%) 1037/1044 (99%)	0.0
AAU68535	Human novel cytokine encoded by cDNA 790CIP2C_6 #1 - Homo sapiens, 1346 aa. [WO200175093-A1, 11-OCT-2001]	1..1044 129..1167	1037/1044 (99%) 1038/1044 (99%)	0.0
AAB93169	Human protein sequence SEQ ID NO:12102 - Homo sapiens, 1014 aa. [EP1074617-A2, 07-FEB-2001]	1..1019 1..1014	1013/1019 (99%) 1013/1019 (99%)	0.0
AAU68534	Human novel cytokine encoded by cDNA 790CIP2C_5 #1 - Homo sapiens, 1324 aa. [WO200175093-A1, 11-OCT-2001]	1..1044 129..1145	1015/1044 (97%) 1015/1044 (97%)	0.0

| ABG27066 | Novel human diagnostic protein #27057 – Homo sapiens, 674 aa. [WO200175067-A2, 11-OCT-2001] | 500..666 47..214 | 166/168 (98%) 166/168 (98%) | 4e-91 |

In a BLAST search of public sequence datbases, the NOV6a protein was found to have homology to the proteins shown in the BLASTP data in Table 6D.

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Table 6D. Public BLASTP Results for NOV6a				
Protein Accession Number	Protein/Organism/Length	NOV6a Residues/ Match Residues	Identities/ Similarities for the Matched Portion	Expect Value
Q9NVE5	CDNA FLJ10785 fis, clone NT2RP4000457, weakly similar to ubiquitin carboxyl-terminal hydrolase 15 (EC 3.1.2.15) – Homo sapiens (Human), 1014 aa (fragment).	1..1019 1..1014	1013/1019 (99%) 1013/1019 (99%)	0.0
Q95KB6	Hypothetical 102.2 kDa protein – Macaca fascicularis (Crab eating macaque) (Cynomolgus monkey), 907 aa (fragment).	143..1024 30..907	844/882 (95%) 860/882 (96%)	0.0
Q8S1J6	Putative ubiquitin carboxyl-terminal hydrolase – Oryza sativa (japonica cultivar-group), 1079 aa.	3..342 223..568	102/359 (28%) 165/359 (45%)	3e-23
Q8VZM4	Putative ubiquitin carboxyl-terminal hydrolase – Arabidopsis thaliana (Mouse-ear cress), 683 aa.	3..202 278..480	72/205 (35%) 105/205 (51%)	3e-23
Q94ED6	Putative ubiquitin carboxyl-terminal hydrolase – Oryza sativa (Rice), 1108 aa.	3..342 273..618	102/359 (28%) 165/359 (45%)	3e-23

PFam analysis predicts that the NOV6a protein contains the domains shown in the Table 6E.

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Table 6E. Domain Analysis of NOV6a			
Pfam Domain	NOV6a Match Region	Identities/ Similarities for the Matched Region	Expect Value
UCH-2	157..354	23/203 (11%) 141/203 (69%)	0.00033

5   **Example 7.**

The NOV7 clone was analyzed, and the nucleotide and encoded polypeptide

sequences are shown in Table 7A.

Table 7A. NOV7 Sequence Analysis			
	SEQ ID NO: 35	692 bp	
NOV7a, CG142564-01 DNA Sequence	GACAGGAGTGAACCCGAGCTGTGCCGACCAACCCCCAGG**ATG**GCGGAAGCTCACCAGGCCGTGGCCTT CCAGTTCACGGTGACCCCAGACGGGGTCGACTTCCGGCTCAGTCGGGAGGCCCTGAAACACGTCTACC TGTCTGGGATCAACTCCTGGAAGAAACGCCTGATCCGCATCAAGAATGGCATCCTCAGGGGCGTGTAC CCTGGCAGCCCCACCAGCTGGCTGGTCGTCATCATGGTAACAGTGGGTTCCTCCTTCTGCAACGTGGA CATCTCCTTGGGGCTGGTCAGTTGCATCCAGAGATGCCTCCCTCAGGGGTGTGGCCCCTACCAGACCC CGCAGACCCGGGCACTTCTCAGCATGGCCATCTTCTCCACGGGCGTCTGGGTGACGGGCATCTTCTTC TTCCGCCAAACCCTGAAGCTGCTTCTCTGCTACCAATCCCAGATCCGCATGTTCGACCCAGAGCAGCA CCCCAATCACCTGGGCGCTGGAGGTGGCTTTGGCCCTGTAGCAGATGATGGCTATGGAGTTTCCTACA TGATTGCAGGCGGAGAACACGATCTTCTTCCACATCTCCAGCAAGTTCTCAAGCTCAGAGACGAACGCC CAGCGCTTTGGAAACCACATCCGCAAAGCCCTGCTGGACATTGCTGATCTTTTTCCAAGTTCCTCAGGC CTACAGC**TGA**AG		
	ORF Start: ATG at 40		ORF Stop: TGA at 688

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	SEQ ID NO: 36	216 aa	MW at 23874.3kD
NOV7a, CG142564-01 Protein Sequence	MAEAHQAVAFQFTVTPDGVDFRLSREALKHVYLSGINSWKKRLIRIKNGILRGVYPGSPTSWLVVIMV TVGSSFCNVDISLGLVSCIQRCLPQGCGPYQTPQTRALLSMAIFSTGVWVTGIFFFRQTLKLLLCYQS QIRMFDPEQHPNHLGAGGGFGPVADDGYGVSYMIAGENTIFFHISSKFSSSETNAQRFGNHIRKALLD IADLFQVPQAYS		

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Further analysis of the NOV7a protein yielded the following properties shown in

Table 7B.

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Table 7B. Protein Sequence Properties NOV7a

PSort analysis:	0.7900 probability located in plasma membrane; 0.6400 probability located in microbody (peroxisome); 0.3000 probability located in Golgi body; 0.2000 probability located in endoplasmic reticulum (membrane)
SignalP analysis:	Cleavage site between residues 5 and 6

A search of the NOV7a protein against the Geneseq database, a proprietary database that contains sequences published in patents and patent publication, yielded several homologous proteins shown in Table 7C.

Table 7C. Geneseq Results for NOV7a				
Geneseq Identifier	Protein/Organism/Length [Patent #, Date]	NOV7a Residues/ Match Residues	Identities/ Similarities for the Matched Region	Expect Value
AAW14438	Type I carnitine palmitoyl transferase-like protein - Homo sapiens, 772 aa. [JP09009969-A, 14-JAN-1997]	1..134 1..134	131/134 (97%) 131/134 (97%)	4e-72
AAE10322	Human carnitine acyltransferase, 26886 - Homo sapiens, 803 aa. [WO200166759-A2, 13-SEP-2001]	1..134 1..132	57/134 (42%) 78/134 (57%)	1e-21
AAY79220	Human transferase TRNSFS-12 - Homo sapiens, 803 aa. [WO200014251-A2, 16-MAR-2000]	1..134 1..132	57/134 (42%) 78/134 (57%)	1e-21
ABB67527	Drosophila melanogaster polypeptide SEQ ID NO 29373 - Drosophila melanogaster, 780 aa. [WO200171042-A2, 27-SEP-2001]	137..210 688..761	43/74 (58%) 55/74 (74%)	6e-19
ABB66942	Drosophila melanogaster polypeptide SEQ ID NO 27618 - Drosophila melanogaster, 782 aa. [WO200171042-A2, 27-SEP-2001]	137..210 690..763	43/74 (58%) 55/74 (74%)	6e-19

In a BLAST search of public sequence datbases, the NOV7a protein was found to have homology to the proteins shown in the BLASTP data in Table 7D.

Table 7D. Public BLASTP Results for NOV7a				
Protein Accession Number	Protein/Organism/Length	NOV7a Residues/ Match Residues	Identities/ Similarities for the Matched Portion	Expect Value
Q9BY90	KIAA1670 protein - Homo sapiens (Human), 598 aa (fragment).	1..134 18..151	133/134 (99%) 133/134 (99%)	2e-73
Q92523	Carnitine O-palmitoyltransferase I, mitochondrial muscle isoform (EC 2.3.1.21) (CPT I) (CPTI-M) (Carnitine palmitoyltransferase I like protein) - Homo sapiens (Human), 772 aa.	1..134 1..134	133/134 (99%) 133/134 (99%)	2e-73
Q924X2	Muscle-type carnitine palmitoyltransferase I (EC 2.3.1.21) (Hypothetical 88.2 kDa protein) - Mus musculus (Mouse), 772 aa.	1..149 1..147	118/149 (79%) 128/149 (85%)	1e-63
O35287	Carnitine palmitoyltransferase I - Mus musculus (Mouse), 772 aa.	1..149 1..147	118/149 (79%) 128/149 (85%)	1e-63
Q9QYP4	Muscle type carnitine palmitoyltransferase I - Mus musculus (Mouse), 772 aa.	1..149 1..147	118/149 (79%) 128/149 (85%)	1e-63

Example 8.

The NOV8 clone was analyzed, and the nucleotide and encoded polypeptide sequences are shown in Table 8A.

Table 8A. NOV8 Sequence Analysis			
	SEQ ID NO: 37	1122 bp	
NOV8a, CG142797-01 DNA Sequence	CTAGATTTTTGAAACATGAATCCTTCACTCCTCCTGGCTGCCTTTTTCCTGGGAATTGCCTCAGCTGC TCTAACACGTGACCACAGTCTAGACGCACAATGGACCAAGTGGAAGGCAAAGCACAAGAGATTATATG ACATGGAGAACATGAAGATGACTGAGCAGCACAATCAGGAATACAGCCAAGGGAAACACAGCTTCACA ATGGCCATGAACACCTTTGGAGACATGACCACTGAAGAATTCAGGCAGGTGATGAATGGTTTTCAATA CCAGAAGCACAGGAACGGGAAACAGTTCCAGGAACGCCTGCTTCTTGAGATCCCCACATCTGTGGACT		

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	GGAGAGAGAAAGGCTACATGACTCCTGTGAAGGATCASGGTCAGTGTGGCTCTTGTTGGGCTTTTAGT GCAACTGGTGCTCTGGAAGGGCAGATGTTCTGGAAAACAGGCAAACTTATCTCACTGAATGAGCAGAA TCTGGTAGACTGCTCTGGGCCTCAAGGCAATGAGGGCTGCAATGGTGGCTTCATGGATAATCCCTTCC GGTATGTTCAGGAGAACGGAGGCCTGGACTCTGAGGCATCCTATCCATATGAAAAAACCTGTAGGTAC AATCCCAAGTATTCTGCTGCTAATGACACTGGCTTTGTGGACATCCCTTCACAGGAGAAGGACCTGGC GAAGGCAGTGGCAACTGTGGGGCCCATCTCTGTTGCTGCTGGTGCAAGCCATGTCTCCTTCCAGTTCT ATAAAAAAGGTATTTATTTTGAGCCACGCTGTGACCCCGAAGGTCTGGATCATGCTATGCTGCTGGTT GGCTACAGCTATGAAGGAGCAGACTCAGATAACAATAAATATTGGCTGGTGAAGAACAGGTATGGTAA AAACTGGGGCATGGATGGCTACATAAAGATGGCCAAAGACCGGAGGAACAACTGTGGAATTGCCACAG CAGCCAGCTACCCCACTGTGTGAGCTGATGGATGGTGATGAGGAAGAACTTGACTGAGGATGGCACAT CCAAAGGAGGAATTTATCTTCAATCTACCAGCCCCTGCTGTGTGGAATGCGCACTTCAATCATTGAAG ATCCAAGTGTGATTGGAATTCTGATATTTTCACA		
	ORF Start: ATG at 16		ORF Stop: TGA at 973

	SEQ ID NO: 38	319 aa	MW at 35984.2kD
NOV8a, CG142797-01 Protein Sequence	MNPSLLLAAFFLGIASAALTRDHSLDAQWTKWKAKHKRLYDMENMKMTEQHNQEYSQGKHSFTMAMNT FGDMTTEEFRQVMNGFQYQKHRNGKQFQERLLLEIPTSVDWREKGYMTPVKDQGQCGSCWAFSATGAL EGQMFWKTGKLISLNEQNLVDCSGPQGNEGCNGGFMDNPFRYVQENGGLDSEASYPYEKTCRYNPKYS AANDTGFVDIPSQEKDLAKAVATVGPISVAAGASHVSFQFYKKGIYFEPRCDPEGLDHAMLLVGYSYE GADSDNNKYWLVKNRYGKNWGMDGYIKMAKDRRNNCGIATAASYPTV		

Further analysis of the NOV8a protein yielded the following properties shown in Table 8B.

Table 8B. Protein Sequence Properties NOV8a	
PSort analysis:	0.8200 probability located in endoplasmic reticulum (membrane); 0.5140 probability located in plasma membrane; 0.2423 probability located in microbody (peroxisome); 0.1000 probability located in endoplasmic reticulum (lumen)
SignalP analysis:	Cleavage site between residues 18 and 19

A search of the NOV8a protein against the Geneseq database, a proprietary database that contains sequences published in patents and patent publication, yielded several homologous proteins shown in Table 8C.

Table 8C. Geneseq Results for NOV8a				
Geneseq Identifier	Protein/Organism/Length [Patent #, Date]	NOV8a Residues/ Match Residues	Identities/ Similarities for the Matched Region	Expect Value

AAU98883	Human protease PRTS1 - Homo sapiens, 334 aa. [WO200238744-A2, 16-MAY-2002]	1..319 1..334	303/334 (90%) 310/334 (92%)	e-180
ABG61771	Novel cathepsin-L precursor-like protein - Homo sapiens, 333 aa. [WO200229058-A2, 11-APR-2002]	1..319 1..333	288/333 (86%) 300/333 (89%)	e-171
ABG66692	Human novel polypeptide #27 - Homo sapiens, 333 aa. [WO200244340-A2, 06-JUN-2002]	1..319 1..333	260/333 (78%) 278/333 (83%)	e-154
ABG66714	Human novel polypeptide #49 - Homo sapiens, 333 aa. [WO200244340-A2, 06-JUN-2002]	1..319 1..333	259/333 (77%) 277/333 (82%)	e-154
ABB77396	Human cathepsin L - Homo sapiens, 333 aa. [DE10050274-A1, 18-APR-2002]	1..319 1..333	249/333 (74%) 274/333 (81%)	e-147

In a BLAST search of public sequence datbases, the NOV8a protein was found to have homology to the proteins shown in the BLASTP data in Table 8D.

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Table 8D. Public BLASTP Results for NOV8a				
Protein Accession Number	Protein/Organism/Length	NOV8a Residues/ Match Residues	Identities/ Similarities for the Matched Portion	Expect Value
P07711	Cathepsin L precursor (EC 3.4.22.15) (Major excreted protein) (MEP) - Homo sapiens (Human), 333 aa.	1..319 1..333	249/333 (74%) 274/333 (81%)	e-147
Q9GKL8	Cysteine protease - Cercopithecus aethiops (Green monkey) (Grivet), 333 aa.	1..319 1..333	247/333 (74%) 273/333 (81%)	e-146
Q9GL24	Cathepsin L (EC 3.4.22.15) - Canis familiaris (Dog), 333 aa.	1..319 1..333	236/334 (70%) 265/334 (78%)	e-138
Q28944	Cathepsin L precursor (EC 3.4.22.15) - Sus scrofa (Pig), 334 aa.	1..319 1..334	228/334 (68%) 263/334 (78%)	e-135

| P25975 | Cathepsin L precursor (EC 3.4.22.15) - Bos taurus (Bovine), 334 aa. | 1..319 1..334 | 222/334 (66%) 261/334 (77%) | e-133 |

PFam analysis predicts that the NOV8a protein contains the domains shown in the Table 8E.

Table 8E. Domain Analysis of NOV8a

Pfam Domain	NOV8a Match Region	Identities/ Similarities for the Matched Region	Expect Value
Peptidase_C1	103..318	123/337 (36%) 194/337 (58%)	2.4e-111

Example 9.

The NOV9 clone was analyzed, and the nucleotide and encoded polypeptide sequences are shown in Table 9A.

Table 9A. NOV9 Sequence Analysis

	SEQ ID NO: 39	1740 bp	
NOV9a, CG143216-01 DNA Sequence	CACGAGGCCGCTAACGGTCCGGCGCCCCTCGGCGTCCGCGCGCCCCCAGCCTGGCGGACGAGCCCGGC GGCGGAGATGGGGGCGACGGGGGCGGCGGGAGCCGCTGCAATCCGTGCTGTGGGTGAAGCAGCAGCGCT GCGCCGTGAGCCTGGAGCCCGCGCGGGCTCTGCTGCGCTGGTGGCGGAGCCCGGGGCCCGGAGCCGGC GCCCCCGGTGCTGATGCCTGCTCTGTGCCTGTATCTGAGATCATCGCCGTTGAGGAAACAGACGTTCA CGGGAAACATCAAGGCAGTGGAAAATGGCAGAAAATGGAAAAGCCTTACGCTTTTACAGTTCACTGTG TAAAGAGAGCACGACGGCACCGCTGGAAGTGGGCGCAGGTGACTTTCTGGTGTCCAGAGGAGCAGCTG TGTCACTTGTGGCTGCAGACCCTGCGGGAGATGCTGGAGAAGCTGACGTCCAGACCAAAGCATTTACT GGTATTTATCAACCCGTTTGGAGGAAAAGGACAAGGCAAGCGGATATATGAAAGAAAAGTGGCACCAC TGTTCACCTTAGCCTCCATCACCACTGACATCATCGTTACTGAACATGCTAATCAGGCCAAGGAGACT CTGTATGAGATTAACATAGACAAATACGACGGCATCGTCTGTGTCGGCGGAGATGGTATGTTCAGCGA GGTGCTGCACGGTCTGATTGGGAGGACGCAGAGGAGCGCCGGGGTCGACCAGAACCACCCCCGGGCTG TGCTGGTCCCCAGTAGCCTCCGGATTGGAATCATTCCCGCAGGGTCAACGGACTGCGTGTGTTACTCC ACCGTGGGCACCAGCGACGCAGAAACCTCGGCGCTGCATATCGTTGTTGGGGACTCGCTGGCCATGGA TGTGTCCTCAGTCCACCACAACAGCACACTCCTTCGCTACTCCGTGTCCCTGCTGGGCTACGGCTTCT ACGGGGACATCATCAAGGACAGTGAGAAGAAACGGTGGTTGGGTCTTGCCAGATACGACTTTTCAGGT TTAAAGACCTTCCTCTCCCACCACTGCTATGAAGGGACAGTGTCCTTCCTCCCTGCACAACACACGGT GGGATCTCCAAGGGATAGGAAGCCCTGCCGGGCAGGATGCTTTGTTTGCAGGCAAAGCAAGCAGCAGC TGGAGGAGGAGCAGAAGAAAGCACTGTATGGTTTGGAAGCTGCGGAGGACGTGGAGGAGTGGCAAGTC GTCTGTGTGGGAAGTTTCTGGCCATCAATGCCACAAACATGTCCTGTGCTTGTCGCCGGAGCCCCAGGGG CCTCTCCCCGGCTGCCCACTTGGGAGACGGGTCTTCTGACCTCATCCTCATCCGGAAATGCTCCAGGT TCAATTTTCTGAGATTTCTCATCAGGCACACCAACCAGCAGGACCAGTTTGACTTCACTTTTGTTGAA GTTTATCGCGTCAAGAAATTCCAGTTTACGTCGAAGCACATGGAGGATGAGGACAGCGACCTCAAGGA GGGGGGGAAGAAGCGCTTTGGGCACATTTGCAGCAGCCACCCCTCCTGCTGCTGCACCGTCTCCAACA GCTCCTGGAACTGCGACGGGGAGGTCCTGCACAGCCCTGCCATCGAGGTCAGAGTCCACTGCCAGCTG GTTCGACTCTTTGCACGAGGAATTGAAGAGAATCCGAAGCCAGACTCACACAGCTGAGAAGCCGGCGT CCTGCTCTCGAACTGGGAAAGTGTGAAAACTATTTAAGAT		
	ORF Start: ATG at 76		ORF Stop: TGA at 1687

	SEQ ID NO: 40	537 aa	MW at 59976.9kD
NOV9a, CG143216-01 Protein Sequence	MGATGAAEPLQSVLWVKQQRCAVSLEPARALLRWWRSPGPGAGAPGADACSVPVSEIIAVEETDVHGK HQGSGKWQKMEKPYAFTVHCVKRARRHRWKWAQVTFWCPEEQLCHLWLQTLREMLEKLTSRPKHLLVF INPFGGKGQGKRIYERKVAPLFTLASITTDIIVTEHANQAKETLYEINIDKYDGIVCVGGDGMFSEVL HGLIGRTQRSAGVDQNHPRAVLVPSSLRIGIIPAGSTDCVCYSTVGTSDAETSALHIVVGDSLAMDVS SVHHNSTLLRYSVSLLGYGFYGDIIKDSEKKRWLGLARYDFSGLKTFLSHHCYEGTVSFLPAQHTVGS PRDRKPCRAGCFVCRQSKQQLEEEQKKALYGLEAAEDVEEWQVVCGKFLAINATNMSCACRRSPRGLS PAAHLGDGSSDLILIRKCSRFNFLRFLIRHTNQQDQFDFTFVEVYRVKKFQFTSKHMEDEDSDLKEGG KKRFGHICSSHPSCCCTVSNSSWNCDGEVLHSPAIEVRVHCQLVRLFARGIEENPKPDSHS

Further analysis of the NOV9a protein yielded the following properties shown in Table 9B.

Table 9B. Protein Sequence Properties NOV9a	
PSort analysis:	0.5121 probability located in microbody (peroxisome); 0.3000 probability located in nucleus; 0.1000 probability located in mitochondrial matrix space; 0.1000 probability located in lysosome (lumen)
SignalP analysis:	No Known Signal Sequence Predicted

A search of the NOV9a protein against the Geneseq database, a proprietary database that contains sequences published in patents and patent publication, yielded several homologous proteins shown in Table 9C.

Table 9C. Geneseq Results for NOV9a				
Geneseq Identifier	Protein/Organism/Length [Patent #, Date]	NOV9a Residues/ Match Residues	Identities/ Similarities for the Matched Region	Expect Value
ABB07857	Human sphingosine kinase-like protein - Homo sapiens, 562 aa. [WO200228906-A2, 11-APR-2002]	1..537 26..562	537/537 (100%) 537/537 (100%)	0.0
ABB07856	Human sphingosine kinase-like protein - Homo sapiens, 537 aa. [WO200228906-A2, 11-APR-2002]	1..537 1..537	537/537 (100%) 537/537 (100%)	0.0

AAM49115	Human ceramide kinase hCERK1 - Homo sapiens, 537 aa. [WO200196575-A1, 20-DEC-2001]	1..537 1..537	535/537 (99%) 536/537 (99%)	0.0
AAY96059	Human sphingosine kinase C - Homo sapiens, 460 aa. [WO200052173-A2, 08-SEP-2000]	78..537 1..460	458/460 (99%) 459/460 (99%)	0.0
AAE07884	Human sphingosine kinase (SphK) protein #2 - Homo sapiens, 471 aa. [WO200160990-A2, 23-AUG-2001]	78..537 1..471	459/471 (97%) 460/471 (97%)	0.0

In a BLAST search of public sequence datbases, the NOV9a protein was found to have homology to the proteins shown in the BLASTP data in Table 9D.

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Table 9D. Public BLASTP Results for NOV9a				
Protein Accession Number	Protein/Organism/Length	NOV9a Residues/ Match Residues	Identities/ Similarities for the Matched Portion	Expect Value
Q8TCT0	Putative lipid kinase - Homo sapiens (Human), 537 aa.	1..537 1..537	537/537 (100%) 537/537 (100%)	0.0
Q9BYB3	KIAA1646 protein - Homo sapiens (Human), 481 aa (fragment).	57..537 1..481	481/481 (100%) 481/481 (100%)	0.0
BAC01155	Ceramide kinases - Mus musculus (Mouse), 531 aa.	1..529 1..529	450/529 (85%) 483/529 (91%)	0.0
Q9UGE5	DA59H18.2 (Novel protein similar to human, mouse, yeast, worm and plant (Predicted) proteins) - Homo sapiens (Human), 326 aa (fragment).	130..444 1..326	314/326 (96%) 315/326 (96%)	0.0
Q9TZI1	T10B11.2 protein - Caenorhabditis elegans, 549 aa.	79..525 115..526	141/458 (30%) 230/458 (49%)	1e-52

PFam analysis predicts that the NOV9a protein contains the domains shown in the

10   Table 9E.

Table 9E. Domain Analysis of NOV9a			
Pfam Domain	NOV9a Match Region	Identities/ Similarities for the Matched Region	Expect Value
PH	32..124	9/93 (10%) 64/93 (69%)	0.38
DAGKc	132..278	32/165 (19%) 100/165 (61%)	0.00015

5   **Example 10.**

The NOV10 clone was analyzed, and the nucleotide and encoded polypeptide sequences are shown in Table 10A.

Table 10A. NOV10 Sequence Analysis			
	SEQ ID NO: 41	772 bp	
NOV10a, CG143787-01 DNA Sequence	AACTGGAGACCACAACTTCATGCTGCGTGGGATCTCCCAACTACCTGCAGTGGCCACCATGTCTTGG GTCCTGCTGCCTGTACTTTGGCTCATTGTTCAAACTCAAGCAATAGCCATAAAGCAAACACCTGAAT TAACGCTCCATGAAATAGTTTGTCCTAAAAAACTTCACATTTTACACAAAAGAGAGATCAAGAACAA CCAGACAGAAAAGCATGGCAAAGAGGAAAGGTATGAACCTGAAGTTCAATATCAGATGATCTTAAAT GGAGAAGAAATCATTCTCTCCCTACAAAAAACCAAGCACCTCCTGGGGCCAGACTACACTGAAACAT TGTACTCACCCAGAGGAGAGGAAATTACCACGAAACCTGAGAACATGGAACACTGTTACTATAAAGG AAACATCCTAAATGAAAAGAATTCTGTTGCCAGCATCAGTACTTGTGACGGGTTGAGAGGATACTTC ACACATCATCACCAAAGATACCTTTTTATCTCAGAAACCAAAGTGCCTGCTGCAAGCACCTATTCCTA CAAATATAATGACAACACCAGTGTGTGGGAACCACCTTCTAGAAGTGGGAGAAGACTGTGATTGTGG CTCTCTTAAGGAGTGTACCAATCTCTGCTGTGAAGCCCTAACGTGTAAACTGAAGCCTGGAACTGAT TGCGGAGGAGATGCTCCAAACCATACCACAGAGTGAATCCAAAAGTCTGCTTCACTGAGATGCTACC TTGCCAGGACAAGAACCAAGAACTCTAACTGTCCC		
	ORF Start: ATG at 20		ORF Stop: TGA at 704

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	SEQ ID NO: 42	228 aa	MW at 25718.4kD
NOV10a, CG143787-01 Protein Sequence	MLRGISQLPAVATMSWVLLPVLWLIVQTQAIAIKQTPELTLHEIVCPKKLHILHKREIKNNQTEKHG KEERYEPEVQYQMILNGEEIILSLQKTKHLLGPDYTETLYSPRGEEITTKPENMEHCYYKGNILNEK NSVASISTCDGLRGYFTHHHQRYLLSQKPKCLLQAPIPTNIMTTPVCGNHLLEVGEDCDCGSLKECT NLCCEALTCKLKPGTDCGGDAPNHTTE		

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	SEQ ID NO: 43	706 bp	
NOV10b, 278889162 DNA Sequence	CACCGGATCCACCATGCTGCGTGGGATCTCCCAACTACCTGCAGTGGCCACCATGTCTTGGGTCCTG CTGCCTGTACTTTGGCTCATTGTTCAAACTCAAGCAATAGCCATAAAGCAAACACCTGAATTAACGC TCCATGAAATAGTTTGTCCTAAAAAACTTCACATTTTACACAAAAGAGAGATCAAGAACAACCAGAC AGAAAAGCATGGCAAAGAGGAAAGGTATGAACCTGAAGTTCAATATCAGATGATCTTAAATGGAGAA		

137

	GAAATCATTCTCTCCCTACAAAAAACCAAGCACCTGCTGGGGGCAGAGTACACTGAAACATTGTACT CACCCAGAGGAGAGGAAATTACCACGAAACCTGAGAACATGGAACACTGTTACTATAAAGGAAACAT CCTAAATGAAAAGAATTCTGTTGCCAGCATCAGTACTTGTGACGGGTTGAGAGGATACTTCACACAT CATCACCAAAGATACCTTTTATCTCAGAAACCAAAGTGCCTGCTGCAAGCACCTATTCCTACAAATA TAATGACAACACCAGTGTGTGGGAACCACCTTCTAGAAGTGGGAGAAGACTGTGATTGTGGCTCTCT TAAGGAGTGTACCAATCTCTGCTGTGAAGCCCTAACGTGTAAACTGAAGCCTGGAACTGATTGCGGA GGAGATGCTCCAAACCATACCACAGAGCTCGAGGGC
	ORF Start: at 2 ORF Stop: end of sequence

	SEQ ID NO: 44	235 aa	MW at 26364.1kD
NOV10b, 278889162 Protein Sequence	TGSTMLRGISQLPAVATMSWVLLPVLWLIVQTQAIAIKQTPELTLHEIVCPKKLHILHKREIKNNQT EKHGKEERYEPEVQYQMILNGEEIILSLQKTKHLLGPDYTETLYSPRGEEITTKPENMEHCYYKGNI LNEKNSVASISTCDGLRGYFTHHHQRYLLSQKPKCLLQAPIPTNIMTTPVCGNHLLEVGEDCDCGSL KECTNLCCEALTCKLKPGTDCGGDAPNHTTELEG		

	SEQ ID NO: 45	118 bp
NOV10c, 278689868 DNA Sequence	CACCGGATCCGAAGTGGGAGAAGACTGTGATTGTGGCTCTCTTAAGGAGTGTACCAATCTCTGCTGT GAAGCCCTAACGTGTAAACTGAAGCCTGGAACTGATTGCGGACTCGAGGGC	
	ORF Start: at 2 ORF Stop: end of sequence	

	SEQ ID NO: 46	39 aa	MW at 3983.4kD
NOV10c, 278689868 Protein Sequence	TGSEVGEDCDCGSLKECTNLCCEALTCKLKPGTDCGLEG		

Sequence comparison of the above protein sequences yields the following sequence relationships shown in Table 10B.

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Table 10B. Comparison of NOV10a against NOV10b and NOV10c.		
Protein Sequence	**NOV10a Residues/ Match Residues**	**Identities/ Similarities for the Matched Region**
NOV10b	1..228 5..232	228/228 (100%) 228/228 (100%)
NOV10c	187..219 4..36	33/33 (100%) 33/33 (100%)

Further analysis of the NOV10a protein yielded the following properties shown in Table 10C.

Table 10C. Protein Sequence Properties NOV10a

PSort analysis:	0.8200 probability located in outside; 0.1900 probability located in lysosome (lumen); 0.1000 probability located in endoplasmic reticulum (membrane); 0.1000 probability located in endoplasmic reticulum (lumen)
SignalP analysis:	Cleavage site between residues 33 and 34

A search of the NOV10a protein against the Geneseq database, a proprietary database that contains sequences published in patents and patent publication, yielded several homologous proteins shown in Table 10D.

Table 10D. Geneseq Results for NOV10a

Geneseq Identifier	Protein/Organism/Length [Patent #, Date]	NOV10a Residues/ Match Residues	Identities/ Similarities for the Matched Region	Expect Value
AAW75769	Human metalloproteinase BS10.55 - Homo sapiens, 470 aa. [WO9839421-A2, 11-SEP-1998]	1..157 1..157	157/157 (100%) 157/157 (100%)	7e-90
AAW28509	Product of clone J5 - Homo sapiens, 470 aa. [WO9707198-A2, 27-FEB-1997]	1..157 1..157	157/157 (100%) 157/157 (100%)	7e-90
AAB53240	Human colon cancer antigen protein sequence SEQ ID NO:780 - Homo sapiens, 110 aa. [WO200055351-A1, 21-SEP-2000]	153..228 35..110	73/76 (96%) 74/76 (97%)	7e-41
ABB11929	Human eMDC II protein homologue, SEQ ID NO:2299 - Homo sapiens, 788 aa. [WO200157188-A2, 09-AUG-2001]	18..159 18..153	71/142 (50%) 99/142 (69%)	2e-32
AAW90865	Human ADAM protein #4 - Homo sapiens, 775 aa. [WO200014227-A1, 16-MAR-2000]	18..159 5..140	71/142 (50%) 99/142 (69%)	2e-32

In a BLAST search of public sequence datbases, the NOV10a protein was found to have homology to the proteins shown in the BLASTP data in Table 10E.

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Table 10E. Public BLASTP Results for NOV10a				
Protein Accession Number	Protein/Organism/Length	NOV10a Residues/ Match Residues	Identities/ Similarities for the Matched Portion	Expect Value
O15204	Disintegrin-protease - Homo sapiens (Human), 470 aa.	1..157 1..157	157/157 (100%) 157/157 (100%)	2e-89
Q9R0X2	Disintegrin metalloprotease precursor - Mus musculus (Mouse), 467 aa.	1..157 1..157	104/157 (66%) 124/157 (78%)	8e-56
Q9XSL6	ADAM 28 precursor (EC 3.4.24.-) (A disintegrin and metalloproteinase domain 28) (eMDC II) - Macaca fascicularis (Crab eating macaque) (Cynomolgus monkey), 776 aa.	14..159 1..141	70/146 (47%) 101/146 (68%)	1e-32
E1262181	SEQUENCE 3 FROM PATENT WO9709430 - unidentified, 530 aa.	18..159 5..140	71/142 (50%) 99/142 (69%)	5e-32
Q9UKQ2	ADAM 28 precursor (EC 3.4.24.-) (A disintegrin and metalloproteinase domain 28) (Metalloproteinase-like, disintegrin-like, and cysteine-rich protein-L) (MDC-L) (eMDC II) (ADAM23) - Homo sapiens (Human), 775 aa.	18..159 5..140	71/142 (50%) 99/142 (69%)	5e-32

PFam analysis predicts that the NOV10a protein contains the domains shown in the Table 10F.

10   Table 10F.

Table 10F. Domain Analysis of NOV10a

Pfam Domain	NOV10a Match Region	Identities/ Similarities for the Matched Region	Expect Value
Pep_M12B_propep	90..201	32/119 (27%) 79/119 (66%)	1.8e-20
disintegrin	187..219	20/33 (61%) 26/33 (79%)	4e-14

Example 11.

The NOV11 clone was analyzed, and the nucleotide and encoded polypeptide

5   sequences are shown in Table 11A.

Table 11A. NOV11 Sequence Analysis			
	SEQ ID NO: 47	484 bp	
NOV11a, CG144112-01 DNA Sequence	ACTGGGTCCGAATCAGTAGGTGACCCCGCCCCTGGATTCTGGAAGACCTCACCATGGGACGCCCCCG ACCTCGTGCGGCCAAGACGTGGATGTTCCTGCTCTTGCTGGGGGGAGCCTGGGCAGGAAATACACAG TACGCCTGGGAGACCACAGCCTACAGAATAAAGATGGCCCAGAAGTGCAGTCCCCGAGAGAATTTTC CTGACACTCTCAACTGTGCAGAAGTAAAAATCTTTCCCCAGAAGAAGTGTGAGGATGCTTACCCGGG GCAGATCACAGATGGCATGGTCTGTGCAGGCAGCAGCAAAGGGGCTGACACGTGCCAGGGCGATTCT GGAGGCCCCCTGGTGTGTGATGGTGCACTCCAGGGCATCACATCCTGGGGCTCAGACCCCTGTGGGA GGTCCGACAAACCTGGCGTCTATACCAACATCTGCCGCTACCTGGACTGGATCAAGAAGATCATAGG CAGCAAGGGC**TGATT**		
	ORF Start: ATG at 54		ORF Stop: TGA at 480

10

	SEQ ID NO: 48	142 aa	MW at 15404.5kD
NOV11a, CG144112-01 Protein Sequence	MGRPRPRAAKTWMFLLLLGGAWAGNTQYAWETTAYRIKMAQKCSPRENFPDTLNCAEVKIFPQKKCE DAYPGQITDGMVCAGSSKGADTCQGDSGGPLVCDGALQGITSWGSDPCGRSDKPGVYTNICRYLDWI KKIIGSKG		

15

	SEQ ID NO: 49	288 bp	
NOV11b, CG144112-04 DNA Sequence	CCCCGCCCC**TGGATTCTGGAAGACCTCACC**ATGGGACGCCCCCGACCTCGTGCGGCCAAGACGTGGA TGTTCCTGCTCTTGCTGGGGGGAGCCTGGGCAGGGCAGGGCGATTCTGGAGGCCCCCTGGTGTGTGA TGGTGCACTCCAGGGCATCACATCCTGGGGCTCAGACCCCTGTGGGAGGTCCGACAAACCTGGCGTC TATACCAACATCTGCCGCTACCTGGACTGGATCAAGAAGATCATAGGCAGCAAGGGC**TGA**TTCTAGG ATAAGCACTAGATCTCCCTT		
	ORF Start: ATG at 31		ORF Stop: TGA at 259

	SEQ ID NO: 50	76 aa	MW at 8110.3kD
NOV11b, CG144112-04 Protein Sequence	MGRPRPRAAKTWMFLLLLGGAWAGQGDSGGPLVCDGALQGITSWGSDPCGRSDKPGVYTNICRYLDW IKKIIGSKG		

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	SEQ ID NO: 51		445 bp	
NOV11c, 255501898 DNA Sequence	CACCAAGCTTATGGGACGCCCCCGACCTCGTGCGGCCAAGACGTGGATGTTCCTGCTCTTGCTGGGG GGAGCCTGGGCAGGAAATACACAGTACGCCTGGGAGACCACAGCCTACAGAATAAAGATGGCCCAGA AGTGCAGTCCCCGAGAGAATTTTCCTGACACTCTCAACTGTGCAGAAGTAAAAATCTTTCCCCAGAA GAAGTGTGAGGATGCTTACCCGGGGCAGATCACAGATGGCATGGTCTGTGCAGGCAGCAGCAAAGGG GCTGACACGTGCCAGGGCGATTCTGGAGGCCCCCTGGTGTGTGATGGTGCACTCCAGGGCATCACAT CCTGGGGCTCAGACCCCTGTGGGAGGTCCGACAAACCTGGCGTCTATACCAACATCTGCCGCTACCT GGACTGGATCAAGAAGATCATAGGCAGCAAGGGCCTCGAGGGC			
	ORF Start: at 2		ORF Stop: end of sequence	

10

	SEQ ID NO: 52	148 aa	MW at 16046.2kD
NOV11c, 255501898 Protein Sequence	TKLMGRPRPRAAKTWMFLLLLGGAWAGNTQYAWETTAYRIKMAQKCSPRENFPDTLNCAEVKIFPQK KCEDAYPGQITDGMVCAGSSKGADTCQGDSGGPLVCDGALQGITSWGSDPCGRSDKPGVYTNICRYL DWIKKIIGSKGLEG		

15

	SEQ ID NO: 53		358 bp	
NOV11d, 255612524 DNA Sequence	CACCAAGCTTGGAAATACACAGTACGCCTGGGAGACCACAGCCTACAGAATAAAGATGGCCCAGAAG TGCAGTCCCCGAGAGAATTTTCCTGACACTCTCAACTGTGCAGAAGTAAAAATCTTTCCCCAGAAGA AGTGTGAGGATGCTTACCCGGGGCAGATCACAGATGGCATGGTCTGTGCAGGCAGCAGCAAAGGGGC TGACACGTGCCAGGGCGATTCTGGAGGCCCCCTGGTGTGTGATGGTGCACTCCAGGGCATCACATCC TGGGGCTCAGACCCCTGTGGGAGGTCCGACAAACCTGGCGTCTATACCAACATCTGCCGCTACCTGG ACTGGATCAAGAAGATCGAGGGC			
	ORF Start: at 2		ORF Stop: end of sequence	

20

	SEQ ID NO: 54	119 aa	MW at 12908.4kD
NOV11d, 255612524 Protein Sequence	TKLGNTQYAWETTAYRIKMAQKCSPRENFPDTLNCAEVKIFPQKKCEDAYPGQITDGMVCAGSSKGA DTCQGDSGGPLVCDGALQGITSWGSDPCGRSDKPGVYTNICRYLDWIKKLEG		

25

	SEQ ID NO: 55	307 bp	
NOV11e, 255612566 DNA Sequence	CACCAAGCTTCAGAAGTGCAGTCCCCGAGAGAATTTTCCTGACACTCTCAACTGTGCAGAAGTAAAA ATCTTTCCCCAGAAGAAGTGTGAGGATGCTTACCCGGGGCAGATCACAGATGGCATGGTCTGTGCAG GCAGCAGCAAAGGGGCTGACACGTGCCAGGGCGATTCTGGAGGCCCCCTGGTGTGTGATGGTGCACT CCAGGGCATCACATCCTGGGGCTCAGACCCCTGTGGGAGGTCCGACAAACCTGGCGTCTATACCAAC ATCTGCCGCTACCTGGACTGGATCAAGAAGCTCGAGGGC		
	ORF Start: at 2	ORF Stop: end of sequence	

	SEQ ID NO: 56	102 aa	MW at 10922.2kD
NOV11e, 255612566 Protein Sequence	TKLQKCSPRENFPDTLNCAEVKIFPQKKCEDAYPGQITDGMVCAGSSKGADTCQGDSGGPLVCDGAL QGITSWGSDPCGRSDKPGVYTNICRYLDWIKKLEG		

	SEQ ID NO: 57	178 bp	
NOV11f, 306434072 DNA Sequence	CACCGGATCCGGGCAGGGCGATTCTGGAGGCCCCCTGGTGTGTGATGGTGCACTCCAGGGCATCACA TCCTGGGGCTCAGACCCCTGTGGGAGGTCCGACAAACCTGGCGTCTATACCAACATCTGCCGCTACC TGGACTGGATCAAGAAGATCATAGGCAGCAAGGGCCTCGAGGGC		
	ORF Start: at 2	ORF Stop: end of sequence	

	SEQ ID NO: 58	59 aa	MW at 6072.7kD
NOV11f, 306434072 Protein Sequence	TGSGQGDSGGPLVCDGALQGITSWGSDPCGRSDKPGVYTNICRYLDWIKKIIGSKGLEG		

	SEQ ID NO: 59	436 bp	
NOV11g, CG144112-02 DNA Sequence	AGTGTGCTGGAATTCGCCCTTACTGGGTCCGAATCAGTAGGTGACCCCGCCCCTGGATTCTTGAAGA CCTCACC**ATG**GGACGCCCCCGACCTCGTGCGGCCAAGACGTGGATGTTCCTGCTCTTGCTGGGGGGA GCCTGGGCAGAGAATTTTCCTGACACTCTCAACTGTGCAGAAGTAAAAATCTTTCCCCAGAAGAAGT GTGAGGATGCTTACCCGGGGCAGATCACAGATGGCATGGTCTGTGCAGGCAGCAGCAAAGGGGCTGA CACGTGCCAGGGCGATTCTGGAGGCCCCCTGGTGTGTGATGGTGCACTCCAGGGCATCACATCCTGG GGCTCAGACCCCTGTGGGAGGTCCGACAAACCTGGCGTCTATACCAACATCTGCCGCTACCTGGACT GGATCAAGAAGATCATAGGCAGCAAGGGC**TGATT**		
	ORF Start: ATG at 75		ORF Stop: TGA at 432

143

	SEQ ID NO: 60	119 aa	MW at 12718.4kD
NOV11g, CG144112-02 Protein Sequence	MGRPRPRAAKTWMFLLLLGGAWAENFPDTLNCAEVKIFPQKKCEDAYPGQITDGMVCAGSSKGADTC QGDSGGPLVCDGALQGITSWGSDPCGRSDKPGVYTNICRYLDWIKKIIGSKG		

5

	SEQ ID NO: 61	845 bp	
NOV11h, CG144112-03 DNA Sequence	CGCCCTTACTGGGTCCGAATCAGTAGGTGACCCCGCCCCTGGATTCTGGAAGACCTCACCATGGGAC GCCCCCGACCTCGTGCGGCCAAGACGTGGATGTTCCTGCTCTTGCTGGGGGGAGCCTGGGCAGGACA CTCCAGGGCACAGGAGGACAAGGTGCTGGGGGGTCATGAGTGCCAACCCCATTCGCAGCCTTGGCAG GCGGCCTTGTTCCAGGGCCAGCAACTACTCTGTGGCGGTGTCCTTGTAGGTGGCAACTGGGTCCTTA CAGCTGCCCACTGTAAAAAACCGAAATACACAGTACGCCTGGGAGACCACAGCCTACAGAATAAAGA TGGCCCAGAGCAAGAAATACCTGTGGTTCAGTCCATCCCACACCCCTGCTACAACAGCAGCGATGTG GAGGACCACAACCATGATCTGATGCTTCTTCAACTGCGTGACCAGGCATCCCTGGGGTCCAAAGTGA AGCCCATCAGCCTGGCAGATCATTGCACCCAGCCTGGCCAGAAGTGCACCGTCTCAGGCTGGGGGCAC TGTCACCAGTCCCCGAGAGAATTTTCCTGACACTCTCAACTGTGCAGAAGTAAAAATCTTTCCCCAG AAGAAGTGTGAGGATGCTTACCCGGGGCAGATCACAGATGGCATGGTCTGTGCAGGCAGCAGCAAAG GGGCTGACACGTGCCAGGGCGATTCTGGAGGCCCCCTGGTGTGTGATGGTGCACTCCAGGGCATCAC ATCCTGGGGCTCAGACCCCTGTGGGAGGTCCGACAAACCTGGCGTCTATACCAACATCTGCCGCTAC CTGGACTGGATCAAGAAGATCATAGGCAGCAAGGGCTGATT
	ORF Start: ATG at 61		ORF Stop: TGA at 841

10

	SEQ ID NO: 62	260 aa	MW at 28047.6kD
NOV11h, CG144112-03 Protein Sequence	MGRPRPRAAKTWMFLLLLGGAWAGHSRAQEDKVLGGHECQPHSQPWQAALFQGQQLLCGGVLVGGNW VLTAAHCKKPKYTVRLGDHSLQNKDGPEQEIPVVQSIPHPCYNSSDVEDHNHDLMLLQLRDQASLGS KVKPISLADHCTQPGQKCTVSGWGTVTSPRENFPDTLNCAEVKIFPQKKCEDAYPGQITDGMVCAGS SKGADTCQGDSGGPLVCDGALQGITSWGSDPCGRSDKPGVYTNICRYLDWIKKIIGSKG		

15

Sequence comparison of the above protein sequences yields the following sequence relationships shown in Table 11B.

Table 11B. Comparison of NOV11a against NOV11b through NOV11h.		
Protein Sequence	NOV11a Residues/ Match Residues	Identities/ Similarities for the Matched Region
NOV11b	97..142 31..76	46/46 (100%) 46/46 (100%)
NOV11c	1..142 4..145	142/142 (100%) 142/142 (100%)
NOV11d	24..139 4..119	114/116 (98%) 115/116 (98%)

NOV11e	41..139 4..102	97/99 (97%) 98/99 (98%)
NOV11f	91..142 5..56	52/52 (100%) 52/52 (100%)
NOV11g	1..142 1..119	119/142 (83%) 119/142 (83%)
NOV11h	44..142 162..260	99/99 (100%) 99/99 (100%)

Further analysis of the NOV11a protein yielded the following properties shown in Table 11C.

Table 11C. Protein Sequence Properties NOV11a	
PSort analysis:	0.3700 probability located in outside; 0.1000 probability located in endoplasmic reticulum (membrane); 0.1000 probability located in endoplasmic reticulum (lumen); 0.1000 probability located in lysosome (lumen)
SignalP analysis:	Cleavage site between residues 24 and 25

A search of the NOV11a protein against the Geneseq database, a proprietary database that contains sequences published in patents and patent publication, yielded several homologous proteins shown in Table 11D.

Table 11D. Geneseq Results for NOV11a				
Geneseq Identifier	Protein/Organism/Length [Patent #, Date]	NOV11a Residues/ Match Residues	Identities/ Similarities for the Matched Region	Expect Value
ABP41332	Human ovarian antigen HCOQP78, SEQ ID NO:2464 - Homo sapiens, 315 aa. [WO200200677-A1, 03-JAN-2002]	44..142 217..315	99/99 (100%) 99/99 (100%)	3e-57
AAU81959	Human PRO322 - Homo sapiens, 260 aa. [WO200109327-A2, 08-FEB-2001]	44..142 162..260	99/99 (100%) 99/99 (100%)	3e-57
ABB84852	Human PRO322 protein sequence SEQ ID NO:72 - Homo sapiens, 260 aa. [WO200200690-A2, 03-JAN-2002]	44..142 162..260	99/99 (100%) 99/99 (100%)	3e-57
ABB95458	Human angiogenesis related protein PRO322 SEQ ID NO: 72 - Homo sapiens, 260 aa. [WO200208284-A2, 31-JAN-2002]	44..142 162..260	99/99 (100%) 99/99 (100%)	3e-57
AAB53087	Human angiogenesis-associated protein PRO322, SEQ ID NO:127 - Homo sapiens, 260 aa. [WO200053753-A2, 14-SEP-2000]	44..142 162..260	99/99 (100%) 99/99 (100%)	3e-57

In a BLAST search of public sequence datbases, the NOV11a protein was found to
have homology to the proteins shown in the BLASTP data in Table 11E.

Table 11E. Public BLASTP Results for NOV11a				
Protein Accession Number	Protein/Organism/Length	NOV11a Residues/ Match Residues	Identities/ Similarities for the Matched Portion	Expect Value
Q9NR68	Serine protease kallikrein/ovasin/neuropsin type 3 - Homo sapiens (Human), 119 aa.	1..142 1..119	119/142 (83%) 119/142 (83%)	9e-66

O60259	Neuropsin precursor (EC 3.4.21.-) (NP) (Kallikrein 8) (Ovasin) (Serine protease TADG-14) (Tumor-associated differentially expressed gene-14 protein) – Homo sapiens (Human), 260 aa.	44..142 162..260	99/99 (100%) 99/99 (100%)	9e-57
O88780	Neuropsin precursor (EC 3.4.21.-) (NP) (Kallikrein 8) (Brain serine protease 1) – Rattus norvegicus (Rat), 260 aa.	38..141 147..259	80/113 (70%) 93/113 (81%)	8e-45
BAB92021	Neuropsin - Mus musculus (Mouse), 176 aa (fragment).	38..141 63..175	81/113 (71%) 92/113 (80%)	1e-44
Q61955	Neuropsin precursor (EC 3.4.21.-) (NP) (Kallikrein 8) - Mus musculus (Mouse), 260 aa.	38..141 147..259	81/113 (71%) 92/113 (80%)	1e-44

PFam analysis predicts that the NOV11a protein contains the domains shown in the Table 11F.

Table 11F. Domain Analysis of NOV11a

Pfam Domain	NOV11a Match Region	Identities/ Similarities for the Matched Region	Expect Value
trypsin	49..134	47/101 (47%) 76/101 (75%)	5.5e-40

Example 12.

The NOV12 clone was analyzed, and the nucleotide and encoded polypeptide sequences are shown in Table 12A.

Table 12A. NOV12 Sequence Analysis			
	SEQ ID NO: 63	1536 bp	
NOV12a, CG144497-01 DNA Sequence	AAGAGCCAAGCCAGCATGTCGGGGACCCGAGCCTCCAACGACCGGCCCCCCGGCGCAGGCGGCGTCA AGCGGGGGCGGCTGCAGCAGGAGGCGGCGGCGACCGGCTCCCGCGTGACGGTGGTGCTGGGCGCGCA GTGGGGGGACGAGGGCAAAGGCAAGGTGGTGGACCTGCTGGCCACGGACGCCGACATCATCAGCCGC TGCCAGGGGGGCAACAACGCCGGCCACACGGTGGTGGTGGATGGGAAAGAGTACGACTTCCACCTGC TGCCCAGCGGCATCATCAACACCAAGGCCGTGTCCTTCATTGGTAACGGGGTGGTCATCCACTTGCC AGGCTTGTTTGAGGAAGCAGAGAAGAATGAAAAGAAAGGTCTGAAGGACTGGGAGAAGAGGCTCATC		

	ATCTCTGACAGAGCCCACCTTGTGTTTGATTTTCACCAGGCTGTCGACGGATTTCAGGAAGTGCAGC		
	GCCAGGCACAAGAGGGGAAGAGTATAGGCACCACCAAGAAGGGAATCGGACCAACCTACTCTTCCAA		
	AGCTGCCCGGACAGGCCTCCGCATCTGCGACCTCCTGTCAGATTTTGATGAGTTTTCCTCCAGATTC		
	AAGAACCTGGCCCACCAGCACCAGTCGATGTTCCCCACCCTGGAAATAGACATTGAAGGCCAACTCA		
	AAAGGCTCAAGGGCTTTGCTGAGCGGATCAGACCCATGGTCCGAGATGGTGTTTACTTTATGTATGA		
	GGCACTCCACGGCCCCCCCAAGAAGATCCTGGTGGAGGGTGCCAACGCCGCCCTCCTCGACATTGAC		
	TTCGGTACCTACCCCTTTGTGACTTCATCCAACTGCACCGTGGGCGGTGTGTGCACGGGCCTGGGCA		
	TCCCCCCGCAGAACATAGGTGACGTGTATGGCGTGGTGAAAGCCTATACCACACGTGTGGGCATCGG		
	GGCCTTCCCCACCGAGCAGATCAACGAGATTGGAGGCCTGCTGCAGACCCGCGGCCACGAGTGGGGA		
	GTGACCACAGGCAGGAAGAGGCGCTGCGGCTGGCTCGACCTGATGATTCTAAGATATGCTCACATGG		
	TCAACGGATTCACTGCGCTGGCCCTGACGAAGCTGGACATCCTGGACGTACTGGGTGAGGTTAAAGT		
	CGGTGTCTCATACAAGCTGAACGGGAAAAGGATTCCCTATTTCCCAGCTAACCAGGAGATGCTTCAG		
	AAGGTCGAAGTTGAGTATGAAACGCTGCCTGGGTGGAAAGCAGACACCACAGGCGCCAGGAGGTGGG		
	AGGACCTGCCCCCACAGGCCCAGAACTACATCCGCTTTGTGGAGAATCACGTGGGAGTCGCAGTCAA		
	ATGGGTTGGTGTTGGCAAGTCAAGAGAGTCGATGATCCAGCTGTTTTAGTCACAGACTGAGCTGATC		
	CCAACAGGCCCTGGCAGCGTCTGGACTTGTGTAAACAGCAGCAGTCACGTTCCTCGGCCGCCACAAC		
	CAACACCAAAGCAGGAAAACCATTTTCTGTACTTTTATATTTCTGTTCAACCTGTTGGTTTC		
ORF Start: ATG at 16		ORF Stop: TAG at 1387	

	SEQ ID NO: 64	457 aa	MW at 50181.0kD
NOV12a, CG144497-01 Protein Sequence	MSGTRASNDRPPGAGGVKRGRLQQEAAATGSRVTVVLGAQWGDEGKGKVVDLLATDADIISRCQGGN NAGHTVVVDGKEYDFHLLPSGIINTKAVSFIGNGVVIHLPGLFEEAEKNEKKGLKDWEKRLIISDRA HLVFDFHQAVDGLQEVQRQAQEGKSIGTTKKGIGPTYSSKAARTGLRICDLLSDFDEFSSRFKNLAH QHQSMFPTLEIDIEGQLKRLKGFAERIRPMVRDGVYFMYEALHGPPKKILVEGANAALLDIDFGTYP FVTSSNCTVGGVCTGLGIPPQNIGDVYGVVKAYTTRVGIGAFPTEQINEIGGLLQTRGHEWGVTTGR KRRCGWLDLMILRYAHMVNGFTALALTKLDILDVLGEVKVGVSYKLNGKRIPYFPANQEMLQKVEVE YETLPGWKADTTGARRWEDLPPQAQNYIRFVENHVGVAVKWVGVGKSRESMIQLF		

Further analysis of the NOV12a protein yielded the following properties shown in Table 12B.

10

Table 12B. Protein Sequence Properties NOV12a	
PSort analysis:	0.5946 probability located in microbody (peroxisome); 0.3000 probability located in nucleus; 0.2377 probability located in lysosome (lumen); 0.1000 probability located in mitochondrial matrix space
SignalP analysis:	No Known Signal Sequence Predicted

A search of the NOV12a protein against the Geneseq database, a proprietary

15   database that contains sequences published in patents and patent publication, yielded

several homologous proteins shown in Table 12C.

Table 12C. Geneseq Results for NOV12a				
Geneseq Identifier	Protein/Organism/Length [Patent #, Date]	NOV12a Residues/	Identities/ Similarities for	Expect Value

		Match Residues	the Matched Region	
AAB41627	Human ORFX ORF1391 polypeptide sequence SEQ ID NO:2782 - Homo sapiens, 314 aa. [WO200058473-A2, 05-OCT-2000]	144..457 1..314	313/314 (99%) 314/314 (99%)	0.0
ABB70971	Drosophila melanogaster polypeptide SEQ ID NO 39705 - Drosophila melanogaster, 447 aa. [WO200171042-A2, 27-SEP-2001]	31..456 24..446	270/427 (63%) 338/427 (78%)	e-161
AAY95049	Candida albicans polypeptide sequence # 17 - Candida albicans, 412 aa. [EP982401-A2, 01-MAR-2000]	35..455 4..409	227/425 (53%) 306/425 (71%)	e-130
AAU23499	Novel human enzyme polypeptide #585 - Homo sapiens, 209 aa. [WO200155301-A2, 02-AUG-2001]	249..457 1..209	208/209 (99%) 209/209 (99%)	e-121
AAW99455	Maize adenylosuccinate synthetase - Zea mays, 484 aa. [US5882869-A, 16-MAR-1999]	24..454 53..482	217/436 (49%) 310/436 (70%)	e-119

In a BLAST search of public sequence datbases, the NOV12a protein was found to have homology to the proteins shown in the BLASTP data in Table 12D.

5

Table 12D. Public BLASTP Results for NOV12a				
Protein Accession Number	Protein/Organism/Length	NOV12a Residues/ Match Residues	Identities/ Similarities for the Matched Portion	Expect Value
BAC04649	CDNA FLJ38602 fis, clone HEART2003836, highly similar to ADENYLOSUCCINATE SYNTHETASE, MUSCLE ISOZYME (EC 6.3.4.4) - Homo sapiens (Human), 457 aa.	1..457 1..457	456/457 (99%) 457/457 (99%)	0.0

P28650	Adenylosuccinate synthetase, muscle isozyme (EC 6.3.4.4) (IMP-- aspartate ligase) (ADSS) (AMPSASE) - Mus musculus (Mouse), 457 aa.	1..457 1..457	441/457 (96%) 453/457 (98%)	0.0
AJMSDS	adenylosuccinate synthase (EC 6.3.4.4), muscle - mouse, 452 aa.	1..425 1..425	411/425 (96%) 421/425 (98%)	0.0
AAH32039	Similar to ADENYLOSUCCINATE SYNTHETASE, MUSCLE ISOZYME (IMP--ASPARTATE LIGASE) (ADSS) (AMPSASE) - Homo sapiens (Human), 502 aa (fragment).	64..457 109..502	392/394 (99%) 394/394 (99%)	0.0
Q9CQL9	Adenylosuccinate synthetase (EC 6.3.4.4) (IMP--aspartate ligase) (ADSS) (AMPSase) - Mus musculus (Mouse), 456 aa.	8..457 4..456	345/453 (76%) 399/453 (87%)	0.0

PFam analysis predicts that the NOV12a protein contains the domains shown in the Table 12E.

Table 12E. Domain Analysis of NOV12a			
Pfam Domain	NOV12a Match Region	Identities/ Similarities for the Matched Region	Expect Value
Ald_Xan_dh_C	396..411	8/16 (50%) 14/16 (88%)	0.43
Adenylsucc_synt	32..455	261/431 (61%) 417/431 (97%)	0

Example 13.

The NOV13 clone was analyzed, and the nucleotide and encoded polypeptide sequences are shown in Table 13A.

Table 13A. NOV13 Sequence Analysis

	SEQ ID NO: 65	278 bp	
NOV13a, CG144686-01 DNA Sequence	TGCCTGTGGGTTTGATTGCTACCACTCTTGCAATTGCTCCTGTCCGCTTTGACAGGGAGAAGGTGTT CCGCGTGAAGCCTCAGGATGAAAAACAAGCAGACATCATAAAGGACTTGGCCAAAACCAGTGAGCTC CGAGATAAAGGCAAATTTGGTTTTCTCCTTCCAGAATCCCGGATAAAGCCAACGTGCAGAGAGACCA TGCTAGCTGTCAAATTTATTGCCAAGTATATCCTCAAGCATACTTCCTAAAGAACTGCCCTCTGTTT GGAATAAGCC		
	ORF Start: at 3		ORF Stop: TAA at 249

	SEQ ID NO: 66	82 aa	MW at 9327.9kD
NOV13a, CG144686-01 Protein Sequence	PVGLIATTLAIAPVRFDREKVFRVKPQDEKQADIIKDLAKTSELRDKGKFGFLLPESRIKPTCRETM LAVKFIAKYILKHTS		

	SEQ ID NO: 67		268 bp
NOV13b, 278690008 DNA Sequence	CACCGGATCCACCCCTGTGGGTTTGATTGCTACCACTCTTGCAATTGCTCCTGTCCGCTTTGACAGG GAGAAGGTGTTCCGCGTGAAGCCTCAGGATGAAAAACAAGCAGACATCATAAAGGACTTGGCCAAAA CCAGTGAGCTCCGAGATAAAGGCAAATTTGGTTTTCTCCTTCCAGAATCCCGGATAAAGCCAACGTG CAGAGAGACCATGCTAGCTGTCAAATTTATTGCCAAGTATATCCTCAAGCATACTTCCCTCGAGGGC		
	ORF Start: at 2		ORF Stop: end of sequence

	SEQ ID NO: 68	89 aa	MW at 9973.6kD
NOV13b, 278690008 Protein Sequence	TGSTPVGLIATTLAIAPVRFDREKVFRVKPQDEKQADIIKDLAKTSELRDKGKFGFLLPESRIKPTC RETMLAVKFIAKYILKHTSLEG		

	SEQ ID NO: 69		94 bp
NOV13c, 278690035 DNA Sequence	CACCGGATCCACCAGTGAGCTCCGAGATAAAGGCAAATTTGGTTTTCTCCTTCCAGAATCCCGGATA AAGCCAACGTGCAGAGAGCTCGAGGGC		
	ORF Start: at 2		ORF Stop: end of sequence

	SEQ ID NO: 70	31 aa	MW at 3452.9kD
NOV13c,	TGSTSELRDKGKFGFLLPESRIKPTCRELEG		

278690035 Protein Sequence		

	SEQ ID NO: 71	1622 bp	
NOV13d, CG144686-02 DNA Sequence	ATGAGGCTCATCCTGCCTGTGGGTTTGATTGCTACCACTCTTGCAATTGCTCCTGTCCGCTTTGACA GGGAGAAGGTGTTCCGCGTGAAGCCCCAGGATGAAAAACAAGCAGACATCATAAAGGACTTGGCCAA AACCAATGAGCTTGACTTCTGGTATCCAGGTGCCACCCACCACGTAGCTGCTAATATGATGGTGGAT TTCCGAGTTAGTGAGAAGGAATCCCAAGCCATCCAGTCTGCCTTGGATCAAAATAAAATGCACTATG AAATCTTGATTCATGATCTACAAGAAGAGATTGAGAAACAGTTTGATGTTAAGGAAGATATCCCAGG CAGGCACAGCTACGCAAAATACAATAATTGGGAAAAGATTGTGGCTTGGACTGAAAAGATGATGGAT AAGTATCCTGAAATGGTCTCTCGTATTAAAATTGGATCTACTGTTGAAGATAATCCACTATATGTTC TGAAGATTGGGGAAAAGAATGAAAGAAGAAAGGCTATTTTTATGGATTGTGGCATTCACGCACGAGA ATGGGTCTCCCCAGCATTCTGCCAGTGGTTTGTCTATCAGGCAACCAAAACTTATGGGAGAAACAAA ATTATGACCAAACTCTTGGACCGAATGAATTTTTACATTCTTCCTGTGTTCAATGTTGATGGATATA TTTGGTCATGGACAAAGAACCGCATGTGGAGAAAAAATCGTTCCAAGAACCAAAACTCCAAATGCAT CGGCACTGACCTCAACAGGAATTTTAATGCTTCATGGAACTCCATTCCTAACACCAATGACCCATGT GCAGATAACTATCGGGGCTCTGCACCAGAGTCCGAGAAAGAGACGAAAGCTGTCACTAATTTCATTA GAAGCCACCTGAATGAAATCAAGGTTTACATCACCTTCCATTCCTACTCCCAGATGCTATTGTTTCC CTATGGATATACATCAAAACTGCCACCTAACCATGAGGACTTGGCCAAAGTTGCAAAGATTGGCACT GATGTTCTATCAACTCGATATGAAACCCGCTACATCTATGGCCCAATAGAATCAACAATTTACCCGA TATCAGGTTCTTCTTTAGACTGGGCTTATGACCTGGGCATCAAACACACATTTGCCTTTGAGCTCCG AGATAAAGGCAAATTTGGTTTTCTCCTTCCAGAATCCCGGATAAAGCCAACGTGCAGAGAGACCATG CTAGCTGTCAAATTTATTGCCAAGTATATCCTCAAGCATACTTCCTAAAGAACTGCCCTCTGTTTGG AATAAGCCAATTAATCCTTTTTTTGTGCCTTTCATCAGAAAGTCAATCTTCAGTTATCCCCAAATGCA GCTTCTATTTCACCTGAATCCTTCTCTTGCTCATTTAAGTCCCATGTTACTGCTGTTTGCTTTTACT TACTTTCAGTAGCACCATAACGAAGTAGCTTTAAGTGAAACCTTTTAACTACCTTTCTTTGCTCCAA GTGAAGTTTGGACCCAGCAGAAAGCATTATTTTGAAAGGTGATATACAGTGGGGCACAGAAAACAAA TGAAAACCCTCAGTTTCTCACAGATTTTCACCATGTGGCTTCATCAATTTATGTGCTAATACAATAA AATAAAATGCACTT
	ORF Start: ATG at 1		ORF Stop: TAA at 1252

	SEQ ID NO: 72	417 aa	MW at 48699.4kD
NOV13d, CG144686-02 Protein Sequence	MRLILPVGLIATTLAIAPVRFDREKVFRVKPQDEKQADIIKDLAKTNELDFWYPGATHHVAANMMVD FRVSEKESQAIQSALDQNKMHYEILIHDLQEEIEKQFDVKEDIPGRHSYAKYNNWEKIVAWTEKMMD KYPEMVSRIKIGSTVEDNPLYVLKIGEKNERRKAIFMDCGIHAREWVSPAFCQWFVYQATKTYGRNK IMTKLLDRMNFYILPVFNVDGYIWSWTKNRMWRKNRSKNQNSKCIGTDLNRNFNASWNSIPNTNDPC ADNYRGSAPESEKETKAVTNFIRSHLNEIKVYITFHSYSQMLLFPYGYTSKLPPNHEDLAKVAKIGT DVLSTRYETRYIYGPIESTIYPISGSSLDWAYDLGIKHTFAFELRDKGKFGFLLPESRIKPTCRETM LAVKFIAKYILKHTS

Sequence comparison of the above protein sequences yields the following sequence relationships shown in Table 13B.

15

Table 13B. Comparison of NOV13a against NOV13b through NOV13d.		
Protein Sequence	NOV13a Residues/ Match Residues	Identities/ Similarities for the Matched Region
NOV13b	1..82 5..86	82/82 (100%) 82/82 (100%)

NOV13c	41..65	25/25 (100%)
	4..28	25/25 (100%)
NOV13d	1..44	43/44 (97%)
	6..49	44/44 (99%)

Further analysis of the NOV13a protein yielded the following properties shown in Table 13C.

Table 13C. Protein Sequence Properties NOV13a	
PSort analysis:	0.5500 probability located in endoplasmic reticulum (membrane); 0.1900 probability located in lysosome (lumen); 0.1000 probability located in endoplasmic reticulum (lumen); 0.1000 probability located in outside
SignalP analysis:	No Known Signal Sequence Predicted

A search of the NOV13a protein against the Geneseq database, a proprietary database that contains sequences published in patents and patent publication, yielded several homologous proteins shown in Table 13D.

Table 13D. Geneseq Results for NOV13a				
Geneseq Identifier	Protein/Organism/Length [Patent #, Date]	NOV13a Residues/ Match Residues	Identities/ Similarities for the Matched Region	Expect Value
AAU84325	Protein CPA3 differentially expressed in breast cancer tissue - Homo sapiens, 417 aa. [WO200210436-A2, 07-FEB-2002]	1..44 6..49	43/44 (97%) 44/44 (99%)	2e-17
AAG75369	Human colon cancer antigen protein SEQ ID NO:6133 - Homo sapiens, 180 aa. [WO200122920-A2, 05-APR-2001]	43..82 141..180	40/40 (100%) 40/40 (100%)	9e-17
AAU04477	Porcine carboxypeptidase B (CpB) protein - Sus scrofa, 306 aa. [WO200151624-A2, 19-JUL-2001]	41..80 266..305	25/40 (62%) 34/40 (84%)	4e-10
AAR75132	Porcine carboxypeptidase B - Sus scrofa. 306 aa.	41..80 266..305	25/40 (62%) 34/40 (84%)	4e-10

	[WO9514096-A1, 26-MAY-1995]				
AAR75131	Porcine Tyr-His-Met Procarboxypeptidase B - Sus scrofa, 404 aa. [WO9514096-A1, 26-MAY-1995]	41..80 364..403	25/40 (62%) 34/40 (84%)	4e-10

In a BLAST search of public sequence datbases, the NOV13a protein was found to have homology to the proteins shown in the BLASTP data in Table 13E.

5

Table 13E. Public BLASTP Results for NOV13a				
Protein Accession Number	Protein/Organism/Length	NOV13a Residues/ Match Residues	Identities/ Similarities for the Matched Portion	Expect Value
P15088	Mast cell carboxypeptidase A precursor (EC 3.4.17.1) (MC-CPA) (Carboxypeptidase A3) - Homo sapiens (Human), 417 aa.	1..44 6..49	43/44 (97%) 44/44 (99%)	5e-17
P97597	Mast cell carboxypeptidase A precursor - Rattus norvegicus (Rat), 412 aa (fragment).	43..82 373..412	37/40 (92%) 39/40 (97%)	1e-14
P21961	Mast cell carboxypeptidase (EC 3.4.17.1) (RMC-CP) (Carboxypeptidase A3) - Rattus norvegicus (Rat), 309 aa.	43..82 270..309	37/40 (92%) 39/40 (97%)	1e-14
P15089	Mast cell carboxypeptidase A precursor (EC 3.4.17.1) (MC-CPA) (Carboxypeptidase A3) - Mus musculus (Mouse), 417 aa.	43..82 378..417	36/40 (90%) 39/40 (97%)	7e-14
P00732	Carboxypeptidase B (EC 3.4.17.2) - Bos taurus (Bovine), 306 aa.	41..80 266..305	26/40 (65%) 36/40 (90%)	7e-11

PFam analysis predicts that the NOV13a protein contains the domains shown in the
10   Table 13F.

Table 13F. Domain Analysis of NOV13a			
Pfam Domain	NOV13a Match Region	Identities/ Similarities for the Matched Region	Expect Value
Zn_carbOpept	41..65	16/30 (53%) 24/30 (80%)	5.6e-08

Example 14.

The NOV14 clone was analyzed, and the nucleotide and encoded polypeptide sequences are shown in Table 14A.

Table 14A. NOV14 Sequence Analysis			
	SEQ ID NO: 73	829 bp	
NOV14a, CG144906-01 DNA Sequence	GCCCTTCGCGGGAGAGGAGGCCATGGGCGCGCGCGGGGCGCTGCTGCTGGCGCTGCTGCTGGCTCGG GCTGGACTCAGGAAGCCGGAGTCGCAGGAGGCGGCGCCCTTATCAGGACCATGCGGCCGACGGGTCA TCACGTCGCGCATCGTGGGTGGAGAGGACGCCGAACTCGGGCGTTGGCCGTGGCAGGGGAGCCTGCG CCTGTGGGATTCCCACGTATGCGGAGTGAGCCTGCTCAGCCACCGCTGGGCACTCACGGCGGCGCAC TGCTTTGAAACCTATAGTGACCTTAGTGATCCCTCCGGGTGGATGGTCCAGTTTGGCCAGCTGACTT CCATGCCATCCTCCACATTTGAGTTTGAGAACCGGACAGACTGCTGGGTGACTGGCTGGGGGTACAT CAAAGAGGATGAGGCACTGCCATCTCCCCACACCCTCCAGGAAGTTCAGGTCGCCATCATAAACAAC TCTATGTGCAACCACCTCTTCCTCAAGTACAGTTTCCGCAAGGACATCTTTGGAGACATGGTTTGTG CTGGCAATGCCCAAGGCGGGAAGGATGCCTGCTTCGGTGACTCAGGTGGACCCTTGGCCTGTAACAA GAATGGACTGTGGTATCAGATTGGAGTCGTGAGCTGGGGAGTGGGCTGTGGTCGGCCCAATCGGCCC GGTGTCTACACCAATATCAGCCACCACTTTGAGTGGATCCAGAAGCTGATGGCCCAGAGTGGCATGT CCCAGCCAGACCCCTCCTGGCCACTACTCTTTTTCCCTCTTCTCTGGGCTCTCCCACTCCTGGGGCC GGTCTGAGCCTACCTGAGCCCATGC		
	ORF Start: ATG at 23		ORF Stop: TGA at 809

	SEQ ID NO: 74	262 aa	MW at 28826.7kD
NOV14a, CG144906-01 Protein Sequence	MGARGALLLALLLARAGLRKPESQEAAPLSGPCGRRVITSRIVGGEDAELGRWPWQGSLRLWDSHVC GVSLLSHRWALTAAHCFETYSDLSDPSGWMVQFGQLTSMPSSTFEFENRTDCWVTGWGYIKEDEALP SPHTLQEVQVAIINNSMCNHLFLKYSFRKDIFGDMVCAGNAQGGKDACFGDSGGPLACNKNGLWYQI GVVSWGVGCGRPNRPGVYTNISHHFEWIQKLMAQSGMSQPDPSWPLLFFPLLWALPLLGPV		

	SEQ ID NO: 75	989 bp	
NOV14b, CG144906-02 DNA Sequence	AATCGCCCTTCGCGGGAGAGGAGGCCATGGGCGCGCGCGGGGCGCTGCTGCTGGCGCTGCTGCTGGC TCGGGCTGGACTCAGGAAGCCGGAGTCGCAGGAGGCGGCGCCCTTATCAGGACCATGCGGCCGACGG GTCATCACGTCGCGCATCGTGGGTGGAGAGGACGCCGAACTCGGGCGTTGGCCGTGGCAGGGGAGCC TGCGCCTGTGGGATTCCCACGTATGCGGAGTGAGCCTGCTCAGCCACCGCTGGGCACTCACGGCGGC GCACTGCTTTGAAACCTATAGTGACCTTAGTGATCCCTCCGGGTGGATGGTCCAGTTTGGCCAGCTG		

| | ACTTCCATGCCATCCTTCTGGAGCCTGCAGGCCTACTACACCCGTTACTTCGTATCGAATATCTATC
TGAGCCCTCGCTACCTGGGGAATTCACCCTATGACATTGCCTTGGTGAAGCTGTCTGCACCTGTCAC
CTACACTAAACACATCCAGCCCATCTGTCTCCAGGCCTCCACATTTGAGTTTGAGAACCGGACAGAC
TGCTGGGTGACTGGCTGGGGGTACATCAAAGAGGATGAGGCACTGCCATCTCCCCACACCCTCCAGG
AAGTTCAGGTCGCCATCATAAACAACTCTATGTGCAACCACCTCTTCCTCAAGTACAGTTTCCGCAA
GGACATCTTTGGAGACATGGTTTGTGCTGGCAATGCCCAAGGCGGGAAGGATGCCTGCTTCGGTGAC
TCAGGTGGACCCTTGGCCTGTAACAGGAATGGACTGTGGTATCAGATTGGAGTCGTGAGCTGGGGAG
TGGGCTGTGGTCGGCCCAATCGGCCCGGTGTCTACACCAATATCAGCCACCACTTTGAGTGGATCCA
GAAGCTGATGGCCCAGAGTGGCATGTCCCAGCCAGACCCCTCCTGGCCACTACTCTTTTTCCCTCTT
CTCTGGGCTCTCCCACTCCTGGGGCCGGTC**TGA**GCCTACCTTAGCCCATGC |
| ORF Start: ATG at 27 | | ORF Stop: TGA at 969 |

	SEQ ID NO: 76	314 aa	MW at 34911.6kD
NOV14b, CG144906-02 Protein Sequence	MGARGALLLALLLARAGLRKPESQEAAPLSGPCGRRVITSRIVGGEDAELGRWPWQGSLRLWDSHVC GVSLLSHRWALTAAHCFETYSDLSDPSGWMVQFGQLTSMPSFWSLQAYYTRYFVSNIYLSPRYLGNS PYDIALVKLSAPVTYTKHIQPICLQASTFEFENRTDCWVTGWGYIKEDEALPSPHTLQEVQVAIINN SMCNHLFLKYSFRKDIFGDMVCAGNAQGGKDACFGDSGGPLACNRNGLWYQIGVVSWGVGCGRPNRP GVYTNISHHFEWIQKLMAQSGMSQPDPSWPLLFFPLLWALPLLGPV		

Sequence comparison of the above protein sequences yields the following sequence relationships shown in Table 14B.

10

Table 14B. Comparison of NOV14a against NOV14b.		
Protein Sequence	**NOV14a Residues/ Match Residues**	**Identities/ Similarities for the Matched Region**
NOV14b	20..240 20..292	219/273 (80%) 221/273 (80%)

Further analysis of the NOV14a protein yielded the following properties shown in
15   Table 14C.

Table 14C. Protein Sequence Properties NOV14a	
PSort analysis:	0.5422 probability located in outside; 0.4639 probability located in lysosome (lumen); 0.2779 probability located in microbody (peroxisome); 0.1900 probability located in plasma membrane
SignalP analysis:	Cleavage site between residues 20 and 21

A search of the NOV14a protein against the Geneseq database, a proprietary database that contains sequences published in patents and patent publication, yielded several homologous proteins shown in Table 14D.

5

Table 14D. Geneseq Results for NOV14a				
Geneseq Identifier	Protein/Organism/Length [Patent #, Date]	NOV14a Residues/ Match Residues	Identities/ Similarities for the Matched Region	Expect Value
AAE17010	Human eosinophil serine protease-1 (esp-1) like enzyme #2 - Homo sapiens, 314 aa. [WO200198503-A2, 27-DEC-2001]	1..262 1..314	262/314 (83%) 262/314 (83%)	e-154
AAB80256	Human PRO303 protein - Homo sapiens, 314 aa. [WO200104311-A1, 18-JAN-2001]	1..262 1..314	262/314 (83%) 262/314 (83%)	e-154
AAU01569	Human secreted protein immunogenic epitope encoded by gene #9 - Homo sapiens, 315 aa. [WO200123547-A1, 05-APR-2001]	1..262 1..314	262/314 (83%) 262/314 (83%)	e-154
AAU02223	Human extracellular serine protease TADG-16 - Homo sapiens, 314 aa. [WO200127257-A1, 19-APR-2001]	1..262 1..314	262/314 (83%) 262/314 (83%)	e-154
AAY91871	Human cancer-specific gene protein, Pro104 - Homo sapiens, 327 aa. [WO200016805-A1, 30-MAR-2000]	1..262 14..327	262/314 (83%) 262/314 (83%)	e-154

In a BLAST search of public sequence datbases, the NOV14a protein was found to have homology to the proteins shown in the BLASTP data in Table 14E.

10

Table 14E. Public BLASTP Results for NOV14a

Protein Accession Number	Protein/Organism/Length	NOV14a Residues/ Match Residues	Identities/ Similarities for the Matched Portion	Expect Value
Q9Y6M0	Testisin precursor (EC 3.4.21.-) (Eosinophil serine protease 1) (ESP- 1) - Homo sapiens (Human), 314 aa.	1..262 1..314	262/314 (83%) 262/314 (83%)	e-154
Q9JHJ7	Testisin precursor (EC 3.4.21.-) (Tryptase 4) - Mus musculus (Mouse), 324 aa.	1..261 1..323	179/326 (54%) 210/326 (63%)	1e-98
Q920S2	Testis serine protease-1 - Mus musculus (Mouse), 322 aa.	1..261 1..321	150/325 (46%) 180/325 (55%)	2e-69
Q9D4I3	4931440B09Rik protein - Mus musculus (Mouse), 282 aa.	32..261 2..281	135/283 (47%) 161/283 (56%)	1e-66
Q9PVX7	Epidermis specific serine protease - Xenopus laevis (African clawed frog), 389 aa.	33..244 17..277	100/264 (37%) 136/264 (50%)	3e-45

PFam analysis predicts that the NOV14a protein contains the domains shown in the Table 14F.

Table 14F. Domain Analysis of NOV14a			
Pfam Domain	NOV14a Match Region	Identities/ Similarities for the Matched Region	Expect Value
trypsin	42..85	24/51 (47%) 36/51 (71%)	2.3e-13
trypsin	119..229	52/121 (43%) 92/121 (76%)	9e-43

Example 15.

The NOV15 clone was analyzed, and the nucleotide and encoded polypeptide sequences are shown in Table 15A.

Table 15A. NOV15 Sequence Analysis			
	SEQ ID NO: 77	716 bp	

NOV15a, CG144997-01 DNA Sequence	GAGTGAGCGATGAGCTGGTTTCTGTTCCTGGCCCACAGAGTGGCCTTGGCCGCCTTGCCCTGCCGCC GCGGCTCTCGCGGGTTCGGGATGTTCTATGCCGTGAGGAGGGGCCGCAAGACCGGGGTCTTTCTGAC CTGGAATGAGTGCAGAGACACGTTTTCCTACATGGGAGACTTCGTCGTCGTCTACACTGATGGCTGC TGCTCCAGTAATGGGCGTAGAAGGCCGCGAGCAGGAATCGGCGTTTACTGGGGGCCGGGCCATCCTT TAAATGTAGGCATTAGACTTCCTGGGCGGCAGACAAACCAAAGAGCGGAAATTCATGCAGCCTGCAA AGCCATTGAACAAGCAAAGACTCAAAACATCAATAAACTGGTTCTGTATACAGACAGTATGTTTACG ATAAATGGTATAACTAACTGGGTTCAAGGTTGGAAGAAAAATGGGTGGAAGACAAGTGCAGGGAAAG AGGTGATCAACAAAGAGGACTTTGTGGCACTGGAGAGGCTTACCCAGGGGATGGACATTCAGTGGAT GCATGTTCCTGGTCATTCGGGATTTATAGGCAATGAAGAAGCTGACAGATTAGCCAGAGAAGGAGCT AAACAATCGGAAGACTGAGCCATGTGACTTTAGTCCTTGGGAGAACTTGAGCCAGCGGCTGTCTTGC TGCCTGTACTTACTGGTGTGGAAAAATAGCCTGCAGGTAGGACCATT
	ORF Start: ATG at 10			ORF Stop: TGA at 619

	SEQ ID NO: 78	203 aa	MW at 22889.0kD
NOV15a, CG144997-01 Protein Sequence	MSWFLFLAHRVALAALPCRRGSRGFGMFYAVRRGRKTGVFLTWNECRDTFSYMGDFVVVYTDGCCSS NGRRRPRAGIGVYWGPGHPLNVGIRLPGRQTNQRAEIHAACKAIEQAKTQNINKLVLYTDSMFTING ITNWVQGWKKNGWKTSAGKEVINKEDFVALERLTQGMDIQWMHVPGHSGFIGNEEADRLAREGAKQS ED		

	SEQ ID NO: 79	631 bp	
NOV15b, 278693648 DNA Sequence	CACCGGATCCACCATGAGCTGGTTTCTGTTCCTGGCCCACAGAGTCGCCTTGGCCGCCTTGCCCTGC CGCCGCGGCTCTCGCGGGTTCGGGATGTTCTATGCCGTGAGGAGGGGCCGCAAGACCGGGGTCTTTC TGACCTGGAATGAGTGCAGAGACACGTTTTCCTACATGGGAGACTTCGTCGTCGTCTACACTGATGG CTGCTGCTCCAGTAATGGGCGTAGAAGGCCGCGAGCAGGAATCGGCGTTTACTGGGGGCCGGGCCAT CCTTTAAATGTAGGCATTAGACTTCCTGGGCGGCAGACAAACCAAAGAGCGGAAATTCATGCAGCCT GCAAAGCCATTGAACAAGCAAAGACTCAAAACATCAATAAACTGGTTCTGTATACAGACAGTATGTT TACGATAAATGGTATAACTAACTGGGTTCAAGGTTGGAAGAAAAATGGGTGGAAGACAAGTGCAGGG AAAGAGGTGATCAACAAAGAGGACTTTGTGGCACTGGAGAGGCTTACCCAGGGGATGGACATTCAGT GGATGCATGTTCCTGGTCATTCGGGATTTATAGGCAATGAAGAAGCTGACAGATTAGCCAGAGAAGG AGCTAAACAATCGGAAGACCTCGAGGGC		
	ORF Start: at 2		ORF Stop: end of sequence

	SEQ ID NO: 80	210 aa	MW at 23534.6kD
NOV15b, 278693648 Protein Sequence	TGSTMSWFLFLAHRVALAALPCRRGSRGFGMFYAVRRGRKTGVFLTWNECRDTFSYMGDFVVVYTDG CCSSNGRRRPRAGIGVYWGPGHPLNVGIRLPGRQTNQRAEIHAACKAIEQAKTQNINKLVLYTDSMF TINGITNWVQGWKKNGWKTSAGKEVINKEDFVALERLTQGMDIQWMHVPGHSGFIGNEEADRLAREG AKQSEDLEG		

	SEQ ID NO: 81	586 bp	
NOV15c, 278480974 DNA Sequence	CACCGGATCCGCCTTGCCCTGCCGCCGCGGCTCTCGCGGGTTCGGGATGTTCTATGCCGTGAGGAGG GGCCGCAAGACCGGGGTCTTTCTGACCTGGAATGAGTGCAGAGACACGTTTTCCTACATGGGAGACT TCGTCGTCGTCTACACTGATGGCTGCTGCTCCAGTAATGGGCGTAGAAGGCCGCGAGCAGGAATCGG CGTTTACTGGGGGCCGGGCCATCCTTTAAATGTAGGCATTAGACTTCCTGGGCGGCAGACAAACCAA AGAGCGGAAATTCATGCAGCCTGCAAAGCCATTGAACAAGCAAAGACTCAAAACATCAATAAACTGG TTCTGTATACAGACAGTATGTTTACGATAAATGGTATAACTAACTGGGTTCAAGGTTGGAAGAAAAA TGGGTGGAAGACAAGTGCAGGGAAAGAGGTGATCAACAAAGAGGACTTTGTGGCACTGGAGAGGCTT		

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	ACCCAGGGGATGGACATTCAGTGGATGCATGTTCCTGGTCATTCGGGATTTATAGGCAATGAAGAAGT
	CTGACAGATTAGCCAGAGAAGGAGCTAAACAATCGGAAGACCTCGAGGGC
ORF Start: at 2	ORF Stop: end of sequence

	SEQ ID NO: 82	195 aa	MW at 21789.5kD
NOV15c, 278480974 Protein Sequence	TGSALPCRRGSRGFGMFYAVRRGRKTGVFLTWNECRDTFSYMGDFVVVYTDGCCSSNGRRRPRAGIG VYWGPGHPLNVGIRLPGRQTNQRAEIHAACKAIEQAKTQNINKLVLYTDSMFTINGITNWVQGWKKN GWKTSAGKEVINKEDFVALERLTQGMDIQWMHVPGHSGFIGNEEADRLAREGAKQSEDLEG		

	SEQ ID NO: 83	457 bp	
NOV15d, 278498047 DNA Sequence	CACCGGATCCGGAGACTTCGTCGTCGTCTACACTGATGGCTGCTGCTCCAGTAATGGGCGTAGAAGG CCGCGAGCAGGAATCGGCGTTTACTGGGGGCCGGGCCATCCTTTAAATGTAGGCATTAGACTTCCTG GGCGGCAGACAAACCAAAGAGCGGAAATTCATGCAGCCTGCAAAGCCATTGAACAAGCAAAGACTCA AAACATCAATAAACTGGTTCTGTATACAGACAGTATGTTTACGATAAATGGTATAACTAACTGGGTT CAAGGTTGGAAGAAAAATGGGTGGAAGACAAGTGCAGGGAAAGAGGTGATCAACAAAGAGGACTTTG TGGCACTGGAGAGGCTTACCCAGGGGATGGACATTCAGTGGATGCATGTTCCTGGTCATTCGGGATT TATAGGCAATGAAGAAGCTGACAGATTAGCCAGAGAAGGAGCTAAACTCGAGGGC		
ORF Start: at 2		ORF Stop: end of sequence	

	SEQ ID NO: 84	152 aa	MW at 16753.8kD
NOV15d, 278498047 Protein Sequence	TGSGDFVVVYTDGCCSSNGRRRPRAGIGVYWGPGHPLNVGIRLPGRQTNQRAEIHAACKAIEQAKTQ NINKLVLYTDSMFTINGITNWVQGWKKNGWKTSAGKEVINKEDFVALERLTQGMDIQWMHVPGHSGF IGNEEADRLAREGAKLEG		

	SEQ ID NO: 85	965 bp	
NOV15e, CG144997-02 DNA Sequence	GAGTGAGCGATGAGCTGGTTTCTGTTCCTGGCCCACAGAGTCGCCTTGGCCGCCTTGCCCTGCCGCC GCGGCTCTCGCGGGTTCGGGATGTTCTATGCCGTGAGGAGGGGCCGCAAGACCGGGGTCTTTCTGAC CTGGAATGAGTGCAGAGCACAGGTGGACCGGTTTCCTGCTGCCAGATTTAAGAAGTTTGCCACAGAG GATGAGGCCTGGGCCTTTGTCAGGAAATCTGCAAGCCCGGAAGTTTCAGAAGGGCATGAAAATCAAC ATGGACAAGAATCGGAGGCGAAAGCCAGCAAGCGACTCCGTGAGCCACTGGATGGAGATGGACATGA AAGCGCAGAGCCGTATGCAAAGCACATGAAGCCGAGCGTGGAGCCGCCTCCAGTTAGCAGAGAC ACGTTTTCCTACATGGGAGACTTCGTCGTCGTCTACACTGATGGCTGCTGCTCCAGTAATGGGCGTA GAAGGCCGCGAGCAGGAATCGGCGTTTACTGGGGGCCAGGCCATCCTTTAAATGTAGGCATTAGACT TCCTGGGCGGCAGACAAACCAAAGAGCGGAAATTCATGCAGCCTGCAAAGCCATTGAACAAGCAAAG ACTCAAAACATCAATAAACTGGTTCTGTATACAGACAGTATGTTTACGATAAATGGTATAACTAACT GGGTTCAAGGTTGGAAGAAAAATGGGTGGAAGACAAGTGCAGGGAAAGAGGTGATCAACAAAGAGGA CTTTGTGGCACTGGAGAGGCTTACCCAGGGGATGGACATTCAGTGGATGCATGTTCCTGGTCATTCG GGATTTATAGGCAATGAAGAAGCTGACAGATTAGCCAGAGAAGGAGCTAAACAATCGGAAGACTGAG CCATGTGACTTTAGTCCTTGGGAGGAACTTGAGCCAGCGGCTGTCTTGCTGCCTGTACTTACTGGTGT GGAAAATAGCCTGCAGGTAGGACCATT		
ORF Start: ATG at 10		ORF Stop: TGA at 868	

	SEQ ID NO: 86	286 aa	MW at 32098.0kD
NOV15e, CG144997-02 Protein Sequence	MSWFLFLAHRVALAALPCRRGSRGFGMFYAVRRGRKTGVFLTWNECRAQVDRFPAARFKKFATEDEA WAFVRKSASPEVSEGHENQHGQESEAKASKRLREPLDGDGHESAEPYAKHMKPSVEPAPPVSRDTFS YMGDFVVVYTDGCCSSNGRRRPRAGIGVYWGPGHPLNVGIRLPGRQTNQRAEIHAACKAIEQAKTQN INKLVLYTDSMFTINGITNWVQGWKKNGWKTSAGKEVINKEDFVALERLTQGMDIQWMHVPGHSGFI GNEEADRLAREGAKQSED		

Sequence comparison of the above protein sequences yields the following sequence relationships shown in Table 15B.

Table 15B. Comparison of NOV15a against NOV15b through NOV15e.		
Protein Sequence	**NOV15a Residues/ Match Residues**	**Identities/ Similarities for the Matched Region**
NOV15b	1..203 5..207	203/203 (100%) 203/203 (100%)
NOV15c	14..203 3..192	189/190 (99%) 190/190 (99%)
NOV15d	54..199 4..149	146/146 (100%) 146/146 (100%)
NOV15e	47..203 130..286	157/157 (100%) 157/157 (100%)

Further analysis of the NOV15a protein yielded the following properties shown in Table 15C.

Table 15C. Protein Sequence Properties NOV15a	
PSort analysis:	0.3700 probability located in outside; 0.1805 probability located in microbody (peroxisome); 0.1080 probability located in nucleus; 0.1000 probability located in endoplasmic reticulum (membrane)
SignalP analysis:	Cleavage site between residues 15 and 16

A search of the NOV15a protein against the Geneseq database, a proprietary database that contains sequences published in patents and patent publication, yielded several homologous proteins shown in Table 15D.

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Table 15D. Geneseq Results for NOV15a

Geneseq Identifier	Protein/Organism/Length [Patent #, Date]	NOV15a Residues/ Match Residues	Identities/ Similarities for the Matched Region	Expect Value
AAY70235	Human RNA-associated protein-16 (RNAAP-16) - Homo sapiens, 286 aa. [WO200011171-A2, 02-MAR-2000]	47..203 130..286	157/157 (100%) 157/157 (100%)	6e-92
AAB97508	Human type II RNase H protein - Homo sapiens, 286 aa. [WO200123613-A1, 05-APR-2001]	47..203 130..286	156/157 (99%) 157/157 (99%)	1e-91
AAY25094	Human type 2 RNase H protein - Homo sapiens, 286 aa. [WO9928447-A1, 10-JUN-1999]	47..203 130..286	156/157 (99%) 157/157 (99%)	1e-91
ABB83371	Human wild-type RNase H1 - Homo sapiens, 286 aa. [WO200240635-A2, 23-MAY-2002]	47..203 130..286	156/157 (99%) 156/157 (99%)	2e-90
ABB83374	Mutant RNase H1, E186Q - Homo sapiens, 286 aa. [WO200240635-A2, 23-MAY-2002]	47..203 130..286	155/157 (98%) 156/157 (98%)	5e-90

5   In a BLAST search of public sequence datbases, the NOV15a protein was found to have homology to the proteins shown in the BLASTP data in Table 15E.

Table 15E. Public BLASTP Results for NOV15a

Protein Accession Number	Protein/Organism/Length	NOV15a Residues/ Match Residues	Identities/ Similarities for the Matched Portion	Expect Value
O60930	Ribonuclease H1 (EC 3.1.26.4) (RNase H1) (Ribonuclease H type II) - Homo sapiens (Human), 286 aa.	47..203 130..286	157/157 (100%) 157/157 (100%)	2e-91

Q8VCR6	Ribonuclease H1 - Mus musculus (Mouse), 285 aa.	47..203 129..285	139/157 (88%) 150/157 (95%)	5e-83
O70338	Ribonuclease H1 (EC 3.1.26.4) (RNase H1) - Mus musculus (Mouse), 285 aa.	47..203 129..285	139/157 (88%) 150/157 (95%)	5e-83
Q91953	mRNA, complete cds, clone CLFEST65 - Gallus gallus (Chicken), 293 aa.	50..202 140..292	117/153 (76%) 135/153 (87%)	4e-70
Q21024	F59A6.6 protein - Caenorhabditis elegans, 369 aa.	58..199 222..363	65/142 (45%) 93/142 (64%)	3e-32

PFam analysis predicts that the NOV15a protein contains the domains shown in the Table 15F.

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Table 15F. Domain Analysis of NOV15a			
Pfam Domain	NOV15a Match Region	Identities/ Similarities for the Matched Region	Expect Value
rnaseH	54..199	65/176 (37%) 125/176 (71%)	2.8e-54

Example 16.

10   The NOV16 clone was analyzed, and the nucleotide and encoded polypeptide sequences are shown in Table 16A.

Table 16A. NOV16 Sequence Analysis			
	SEQ ID NO: 87	2274 bp	
NOV16a, CG145494-01 DNA Sequence	CCCCTAGTGACACTCAGGAAATGCTTGTCTCCGGCTGTTAAGGAATAATTTCAGAGTACTATGGATC ATGCTGAAGAAAATGAAATCCTTGCAGCAACCCAGAGGTACTATGTGGAAAGGCCTATCTTTAGTCA TCCGGTCCTCCAGGAAAGACTACACACAAAGGACAAGGTTCCTGATTCCATTGCGGATAAGCTGAAA CAGGCATTCACATGTACTCCTAAAAAAATAAGAAATATCATTTATATGTTCCTACCCATAACTAAAT GGCTGCCAGCATACAAATTCAAGGAATATGTGTTGGGTGACTTGGTCTCAGGCATAAGCACAGGGGT GCTTCAGCTTCCTCAAGGCTTAGCCTTTGCAATGCTGGCAGCTGTGCCTCCAATATTTGGCCTGTAC TCTTCATTTTACCCTGTTATCATGTATTGTTTTCTTGGAACCTCCAGCACACATATCCATAGGTCCTT TTGCTGTTATTAGCCTGATGATTGGTGGTGTAGCTGTTCGATTAGTACCAGATGATATAGTCATTCC AGGAGGAGTAAATGCAACCAATGGCACAGAGGCCAGAGATGCCTTGAGAGTGAAAGTCGCCATGTCT GTGACCTTACTTTCAGGAATCATTCAGTTTTGCCTAGGTGTCTGTAGGTTTGGATTTGTGGCCATAT ATCTCACAGAGCCTCTGGTCCGTGGGTTTACCACCGCAGCAGCTGTGCATGTCTTCACCTCCATGTT AAAATATCTGTTTGGAGTTAAAACAAAGCGGTACAGTGGAATCTTTTCCGTGGTGTATAGTACAGTT GCTGTGTTGCAGAATGTTAAAAACCTCAACGTGTGTTCCCTAGGCGTCGGGCTGATGGTTTTTGGTT TGCTGTTGGGTGGCAAGGAGTTTAATGAGAGATTTAAAGAGAAATTGCCGGCCGCCTATTCCTTTAGA GTTCTTTGCGGTCGTAATGGGAACTGGCATTTCAGCTGGGTTTAACTTGAAAGAATCATACAATGTG GATGTCGTTGGAACACTTCCTCTAGGGCTGCTACCTCCAGCCAATCCGGACACCAGCCTCTTCCACC TTGTGTACGTAGATGCCATTGCCATAGCCATCGTTGGATTTTCAGTGACCATCTCCATGGCCAAGAC

	CTTAGCAAATAAACATGGCTACCAGGTTGACGGCAA(...)AGGAG(...)(...)GGGA(...)(...)(...)
	TCCATTGGCTCACTCTTCCAGACCTTTTCAATTTCATGCTCCTTGTCTCGAAGCCTTGTTCAGGAGG
	GAACCGGTGGGAAGACACAGGCTGTGCTGTCGGCCATTGTGATTGTCAACCTGAAGGGAATGTTTAT
	GCAGTTCTCAGATCTCCCCTTTTTCTGGAGAACCAGCAAAATAGAGCTGACCATCTGGCTTACCACT
	TTTGTGTCCTCCTTGTTCCTGGGATTGGACTATGGTTTGATCACTGCTGTGATCATTGCTCTGCTGA
	CTGTGATTTACAGAACACAGAGTCCAAGCTACAAAGTCCTTGGAAAGCTTCCTGAAACTGATGTGTA
	TATTGATATAGACGCATATGAGGAGGTGAAAGAAATTCCTGGAATAAAAATATTTCAAATAAATGCA
	CCAATTTACTATGCAAATAGCGACTTGTATAGCAATGCATTAAAACGAAAGACTGGAGTGAACCCAG
	CAGTCATCATGGGAGCAAGGAGAAAGGCCATGCGGAAGTACGCTAAGGAAGTCGGAAATGCAAATAT
	GGCCAACGCAACTGTTGTCAAAGCAGATGCAGAAGTAGATGGAGAGGATGCTACCAAGCCTGAAGAA
	GAGGATGGTGAAGTAAAATATCCCCCAATAGTGATCAAAAGCACATTTCCTGAGGAAATGCAAAGAT
	TTATGCCCCCAGGGGATAACGTCCACACTGTCATTTTGGATTTCACTCAAGTCAATTTTATTGATTC
	TGTTGGAGTGAAAACTCTGGCAGGGATTGTAAAAGAATATGGAGACGTCGGTATATATGTATACTTA
	GCAGGATGCAGTGCACAAGTTGTGAATGACCTCACTCGGAATAGATTTTTTGAAAATCCTGCCCTAT
	GGGAGCTGCTGTTCCACAGCATTCATGATGCAGTTTTAGGCAGCCAACTTAGAGAGGCACTTGCTGA
	ACAGGAAGCCTCGGCTCCCCCTTCCCAGGAGGACTTGGAGCCCAATGCCACTCCTGCCACTCCTGAG
	GCA**TAG**ATGAGGACCTCACCCTAGGATGGGGTTATAAGCCTCTCATGAAGTTCATAATTTACA
ORF Start: ATG at 61		**ORF Stop: TAG at 2215**

	SEQ ID NO: 88	**718 aa**	**MW at 78546.4kD**
NOV16a, CG145494-01 Protein Sequence	MDHAEENEILAATQRYYVERPIFSHPVLQERLHTKDKVPDSIADKLKQAFTCTPKKIRNIIYMFLPI TKWLPAYKFKEYVLGDLVSGISTGVLQLPQGLAFAMLAAVPPIFGLYSSFYPVIMYCFLGTSRHISI GPFAVISLMIGGVAVRLVPDDIVIPGGVNATNGTEARDALRVKVAMSVTLLSGIIQFCLGVCRFGFV AIYLTEPLVRGFTTAAAVHVFTSMLKYLFGVKTKRYSGIFSVVYSTVAVLQNVKNLNVCSLGVGLMV FGLLLGGKEFNERFKEKLPAPIPLEFFAVVMGTGISAGFNLKESYNVDVVGTLPLGLLPPANPDTSL FHLVYVDAIAIAIVGFSVTISMAKTLANKHGYQVDGNQELIALGLCNSIGSLFQTFSISCSLSRSLV QEGTGGKTQAVLSAIVIVNLKGMFMQFSDLPFFWRTSKIELTIWLTTFVSSLFLGLDYGLITAVIIA LLTVIYRTQSPSYKVLGKLPETDVYIDIDAYEEVKEIPGIKIFQINAPIYYANSDLYSNALKRKTGV NPAVIMGARRKAMRKYAKEVGNANMANATVVKADAEVDGEDATKPEEEDGEVKYPPIVIKSTPPEEM QRFMPPGDNVHTVILDFTQVNFIDSVGVKTLAGIVKEYGDVGIYVYLAGCSAQVVNDLTRNRFFENP ALWELLFHSIHDAVLGSQLREALAEQEASAPPSQEDLEPNATPATPEA

Further analysis of the NOV16a protein yielded the following properties shown in Table 16B.

Table 16B. Protein Sequence Properties NOV16a	
PSort analysis:	0.6000 probability located in plasma membrane; 0.4000 probability located in Golgi body; 0.3200 probability located in nucleus; 0.3000 probability located in endoplasmic reticulum (membrane)
SignalP analysis:	No Known Signal Sequence Predicted

A search of the NOV16a protein against the Geneseq database, a proprietary database that contains sequences published in patents and patent publication, yielded several homologous proteins shown in Table 16C.

Table 16C. Geneseq Results for NOV16a

Geneseq Identifier	Protein/Organism/Length [Patent #, Date]	NOV16a Residues/ Match Residues	Identities/ Similarities for the Matched Region	Expect Value
AAY71067	Human membrane transport protein, MTRP-12 - Homo sapiens, 758 aa. [WO200026245-A2, 11-MAY-2000]	9..684 15..738	291/741 (39%) 433/741 (58%)	e-148
AAG67162	Amino acid sequence of a human 32613 transporter polypeptide - Homo sapiens, 751 aa. [WO200164875-A2, 07-SEP-2001]	9..684 15..731	289/734 (39%) 432/734 (58%)	e-147
ABG61914	Prostate cancer-associated protein #115 - Mammalia, 790 aa. [WO200230268-A2, 18-APR-2002]	16..699 20..741	268/723 (37%) 419/723 (57%)	e-144
AAM51696	Human pendrin SEQ ID NO 2 - Homo sapiens, 780 aa. [JP2001228146-A, 24-AUG-2001]	16..699 20..741	268/723 (37%) 419/723 (57%)	e-144
AAM51695	Mouse pendrin SEQ ID NO 1 - Mus sp, 780 aa. [JP2001228146-A, 24-AUG-2001]	16..688 20..730	270/713 (37%) 414/713 (57%)	e-142

In a BLAST search of public sequence datbases, the NOV16a protein was found to have homology to the proteins shown in the BLASTP data in Table 16D.

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Table 16D. Public BLASTP Results for NOV16a				
Protein Accession Number	Protein/Organism/Length	NOV16a Residues/ Match Residues	Identities/ Similarities for the Matched Portion	Expect Value
P58743	Prestin - Homo sapiens (Human), 744 aa.	1..718 1..744	718/744 (96%) 718/744 (96%)	0.0
Q9JKQ2	Prestin - Meriones unguiculatus (Mongolian jird) (Mongolian gerbil), 744 aa.	1..718 1..744	679/744 (91%) 699/744 (93%)	0.0
Q99NH7	Prestin - Mus musculus (Mouse), 744 aa.	1..718 1..744	680/744 (91%) 700/744 (93%)	0.0

| Q9EPH0 | Prestin - Rattus norvegicus (Rat), 744 aa. | 1..718
1..744 | 677/744 (90%)
699/744 (92%) | 0.0 |
| AAH28856 | Solute carrier family 26, member 6 - Mus musculus (Mouse), 735 aa. | 16..684
8..715 | 282/718 (39%)
432/718 (59%) | e-148 |

PFam analysis predicts that the NOV16a protein contains the domains shown in the Table 16E.

5

Table 16E. Domain Analysis of NOV16a			
Pfam Domain	**NOV16a Match Region**	**Identities/ Similarities for the Matched Region**	**Expect Value**
COX3	334..458	31/266 (12%) 80/266 (30%)	0.7
Sulfate_transp	193..477	111/328 (34%) 234/328 (71%)	7e-78
STAS	500..683	34/188 (18%) 124/188 (66%)	1.4e-12

Example 17.

10   The NOV17 clone was analyzed, and the nucleotide and encoded polypeptide sequences are shown in Table 17A.

Table 17A. NOV17 Sequence Analysis			
	SEQ ID NO: 89	2124 bp	
NOV17a, CG145722-01 DNA Sequence	AAGCTGAGGTCTTATAGATTGGTGGTACTTAAGGCAGAAAATTAACACCGTGTTTTGTAGCTGTTAG TTGGTAGAGGGAAATTCAGGCTACCGTCGCGAAACCTGCAGGTTAAGTTATTTTCTCCTCCCTGCTT CTGTAGGTTCACAGCGTTCCCTTCTGATAGAGCTTTTTTGTCTGTGTTGTAAAGCTCTTTGGCTGAGA TGGATGACAAAGATATTGACAAAGAACTAAGGCAGAAATTAAACTTTTCCTATTGTGAGGAGACTGA GATTGAAGGGCAGAAGAAAGTAGAAGAAAGCAGGGAGGCTTCGAGCCAAACCCCAGAGAAGGGTGAA GTGCAGGATTCAGAGGCAAAGGGTACACCACCTTGGACTCCCCTTAGCAACGTGCATGAGCTCGACA CATCTTCGGAAAAAGACAAAGAAAGTCCAGATCAGATTTTGAGGACTCCAGTGTCACACCCTCTCAA ATGTCCTGAGACACCAGCCCAACCAGACAGCAGGAGCAAGCTGCTGCCCAGTGACAGCCCCTCTACT CCCAAAACCATGCTGAGCCGGTTGGTGATTTCTCCAACAGGGAAGCTTCCTTCCAGAGGCCCTAAGC ATTTGAAGCTCACACCTGCTCCCCTCAAGGATGAGATGACCTCATTGGCTCTGGTCAATATTAATCC CTTCACTCCAGAGTCCTATAAAAAATTATTTCTTCAATCTGGTGGCAAGAGGAAAATAAGAAGATGT GTTTTACGAGAAACCAACATGGCTTCCCGCTATGAAAAAGAATTCTTGGAGGTTGAAAAAATTGGGG TTGGCGAATTTGGTACAGTCTACAAGTGCATTAAGAGGCTGGATGGATGTGTTTATGCAATAAAGCG CTCTATGAAAACTTTTACAGAATTATCAAATGAGAATTCGGCTTTGCATGAAGTTTATGCTCACGCA GTGCTTGGGCATCACCCCCATGTGGTACGTTACTATTCCTCATGGGCAGAAGATGACCACATGATCA TTCAGAATGAATACTGCAATGGTGGGAGTTTGCAAGCTGCTATATCTGAAAACACTAAGTCTGGCAA TCATTTTGAAGAGCCAAAACTCAAGGACATCCTTCTACAGATTTCCCTTGGCCTTAATTACATCCAC AACTCTAGCATGGTACACCTGGACATCAAACCTAGTAATATATTCATTTGTCACAAGATGCAAAGTG AATCCTCTGGAGTCATAGAAGAAGTTGAAAATGAAGCTGATTGGTTTCTCTCTGCCAATGTGATGTA TAAAATTGGTGACCTGGGCCACGCAACATCAATAAACAAACCCAAAGTGGAAGAAGGAGATAGTCGC

	TTCCTGGCTAATGAGATTTTGCAAGAGGATTACCGC~~AGGTTCCCAAAGCAGACATATTTTCCTTGG~~
	GATTAACAATTGCAGTGGCTGCAGGAGCAGAGTCATTGCCCACCAATGGTGCTGCATGGCACCATAT
	CCGCAAGGGTAACTTTCCGGACGTTCCTCAGGAGCTCTCAGAAAGCTTTTCCAGTCTGCTCAAGAAC
	ATGATCCAACCTGATGCCGAACAGAGACCTTCTGCAGCAGCTCTGGCCAGAAATACAGTTCTCCGGC
	CTTCCCTGGGAAAAACAGAAGAGCTCCAACAGCAGCTGAATTTGGAAAAGTTCAAGACTGCCACACT
	GGAAAGGGAACTGAGAGAGACCCAGCAGGCCCAGTCACCCCAGGGATATACCCATCATGGTGACACT
	GGGGTCTCTGGGACCCACACAGGATCAAGAAGCACAAAACGCCTGGTGGGAGGAAAGAGTGCAAGGT
	CTTCAAGCTTTACCTGTGAG**TAA**TCTTCCCCTTAAGAACTCATTTTGCAGCCGGGCGTGGTGGCTCA
	CGCCTGTAATCCCAACACTTTGGGAGGCCAAGGCAGGTGGATCATGAGGTCAGGAGATCGAAACCAT
	CCTGGCTAACACGGTGAAACCCCATCTCTACTAAAAATACAAAAAATTAGCAGGGCGAGGTGGCAGG
	CGCCTATAATCCCAGCTACTCAGGAGGCTGAGGAAGGAGAATCGCTTGAACCCGGGAGGTGGAGCTT
	GCAGTGAGCTGAGATCACACCACTGCACTCCAGCCTGGGCAACAGAG
ORF Start: ATG at 201			ORF Stop: TAA at 1830

	SEQ ID NO: 90	543 aa	MW at 60514.5kD
NOV17a, CG145722-01 Protein Sequence	MDDKDIDKELRQKLNFSYCEETEIEGQKKVEESREASSQTPEKGEVQDSEAKGTPPWTPLSNVHELD TSSEKDKESPDQILRTPVSHPLKCPETPAQPDSRSKLLPSDSPSTPKTMLSRLVISPTGKLPSRGPK HLKLTPAPLKDEMTSLALVNINPFTPESYKKLFLQSGGKRKIRRCVLRETNMASRYEKEFLEVEKIG VGEFGTVYKCIKRLDGCVYAIKRSMKTFTELSNENSALHEVYAHAVLGHHPHVVRYYSSWAEDDHMI IQNEYCNGGSLQAAISENTKSGNHFEEPKLKDILLQISLGLNYIHNSSMVHLDIKPSNIFICHKMQS ESSGVIEEVENEADWFLSANVMYKIGDLGHATSINKPKVEEGDSRFLANEILQEDYRHLPKADIFAL GLTIAVAAGAESLPTNGAAWHHIRKGNFPDVPQELSESFSSLLKNMIQPDAEQRPSAAALARNTVLR PSLGKTEELQQQLNLEKFKTATLERELREAQQAQSPQGYTHHGDTGVSGTHTGSRSTKRLVGGKSAR SSSFTCE

Further analysis of the NOV17a protein yielded the following properties shown in Table 17B.

Table 17B. Protein Sequence Properties NOV17a

PSort analysis:	0.4500 probability located in cytoplasm; 0.3000 probability located in microbody (peroxisome); 0.1000 probability located in mitochondrial matrix space; 0.1000 probability located in lysosome (lumen)
SignalP analysis:	No Known Signal Sequence Predicted

A search of the NOV17a protein against the Geneseq database, a proprietary database that contains sequences published in patents and patent publication, yielded several homologous proteins shown in Table 17C.

Table 17C. Geneseq Results for NOV17a

Geneseq Identifier	Protein/Organism/Length [Patent #, Date]	NOV17a Residues/ Match Residues	Identities/ Similarities for the Matched Region	Expect Value

AAB62519	Xenopus Wee1 protein catalytic domain (residues 210-443) - Xenopus sp, 240 aa. [US6225101-B1, 01-MAY-2001]	188..431 1..240	170/244 (69%) 191/244 (77%)	1e-94
AAY51401	Xenopus sp. Wee1 catalytic domain protein fragment - Xenopus sp, 240 aa. [US6020194-A, 01-FEB-2000]	188..431 1..240	170/244 (69%) 191/244 (77%)	1e-94
ABB60693	Drosophila melanogaster polypeptide SEQ ID NO 8871 - Drosophila melanogaster, 609 aa. [WO200171042-A2, 27-SEP-2001]	109..501 101..551	180/464 (38%) 257/464 (54%)	9e-78
AAY96776	Z. mays partial wee1 kinase - Zea mays, 525 aa. [WO200037645-A2, 29-JUN-2000]	185..465 264..513	103/282 (36%) 153/282 (53%)	3e-45
AAY96770	Z. mays partial wee1 kinase - Zea mays, 403 aa. [WO200037645-A2, 29-JUN-2000]	185..465 142..391	103/282 (36%) 153/282 (53%)	3e-45

In a BLAST search of public sequence datbases, the NOV17a protein was found to have homology to the proteins shown in the BLASTP data in Table 17D.

5

Table 17D. Public BLASTP Results for NOV17a				
Protein Accession Number	Protein/Organism/Length	NOV17a Residues/ Match Residues	Identities/ Similarities for the Matched Portion	Expect Value
O95017	WUGSC:H_DJ0894A10.2 protein - Homo sapiens (Human), 541 aa (fragment).	1..541 1..541	541/541 (100%) 541/541 (100%)	0.0
P47817	Wee1-like protein kinase (EC 2.7.1.112) - Xenopus laevis (African clawed frog), 555 aa.	10..542 11..552	291/560 (51%) 352/560 (61%)	e-143
O57473	Wee1 homolog - Xenopus laevis (African clawed frog), 554 aa.	10..542 11..551	294/566 (51%) 357/566 (62%)	e-143

| Q8QGV2 | Wee1B kinase - Xenopus laevis (African clawed frog), 595 aa. | 10..541 20..593 | 263/579 (45%) 350/579 (60%) | e-122 |
| Q63802 | Wee1-like protein kinase (EC 2.7.1.112) - Rattus norvegicus (Rat), 646 aa. | 92..541 168..644 | 236/484 (48%) 308/484 (62%) | e-118 |

PFam analysis predicts that the NOV17a protein contains the domains shown in the Table 17E.

Table 17E. Domain Analysis of NOV17a

Pfam Domain	NOV17a Match Region	Identities/ Similarities for the Matched Region	Expect Value
pkinase	194..462	73/310 (24%) 193/310 (62%)	6.4e-45

Example 18.

The NOV18 clone was analyzed, and the nucleotide and encoded polypeptide sequences are shown in Table 18A.

Table 18A. NOV18 Sequence Analysis

	SEQ ID NO: 91	753 bp	
NOV18a, CG145754-01 DNA Sequence	TCCCTTCTCCTGCCCCTGCAGATCCTACTGCTATCCTTAGCCTTGGAAACTGCAGGAGAAGAAGCCC AGGGTGACAAGATTATTGATGGCGCCCCATGTGCAAGAGGCTCCCACCCATGGCAGGTGGCCCTGCT CAGTGGCAATCAGCTCCACTGCGGAGGCGTCCTGGTCAATGAGCGCTGGGTGCTCACTGCCGCCCAC TGCAAGATGAATGAGTACACCGTGCACCTGGGCAGTGATACGCTGGGCGACAGGAGAGCTCAGAGGA TCAAGGCCTCGAAGTCATTCCGCCACCCCGGCTACTCCACACAGACCCATGTTAATGACCTCATGCT CGTGAAGCTCAATAGCCAGGCCAGGCTGTCATCCATGGTGAAGAAAGTCAGGCTGCCCTCCCGCTGC GAACCCCCTGGAACCACCTGTACTGTCTCCGGCTGGGGCACTACCACGAGCCCAGATGTGACCTTTC CCTCTGACCTCATGTGCGTGGATGTCAAGCTCATCTCCCCCCAGGACTGCACGAAGGTTTACAAGGA CTTACTGGAAAATTCCATGCTGTGCGCTGGCATCCCCGACTCCAAGAAAAACGCCTGCAATGGTGAC TCAGGGGGACCGTTGGTGTGTCAGAGGTACCCTGCAAGGTCTGGTGTCCTGGGGAACTTTCCCTTGCG GCCAACCCAATGACCCAGGAGTCTACACTCAAGTGTGCAAGTTCACCAAGTGGATAAATGACACCAT GAAAAAGCATCGCTAA		
	ORF Start: at 1		ORF Stop: TAA at 751

	SEQ ID NO: 92	250 aa	MW at 27166.0kD
NOV18a, CG145754-01	SLLLPLQILLLSLALETAGEEAQGDKIIDGAPCARGSHPWQVALLSGNQLHCGGVLVNERWVLTAAH CKMNEYTVHLGSDTLGDRRAQRIKASKSFRHPGYSTQTHVNDLMLVKLNSQARLSSMVKKVRLPSRC EPPGTTCTVSGWGTTTSPDVTFPSDLMCVDVKLISPODCTKVYKDLLENSMLCAGIPDSKKNACNGD		

Protein Sequence	SGGPLVCRGTLQGLVSWGTFPCGQPNDPGVYTQVCKFTKWINDTMKKHR

	SEQ ID NO: 93	862 bp	
NOV18b, CG145754-03 DNA Sequence	ACTGGGTCCGAATCAGTAGGTGACCCCGCCCCTGGATTCTGGAAGACCTCACC**ATG**GGACGCCCCCG ACCTCGTGCGGCCAAGACGTGGATGTTCCTGCTCTTACTGGGGGGAGCCTGGGCAGCCAGGGGTGAC AAGATTATTGATGGCGCCCCATGTGCAAGAGGCTCCCACCCATGGCAGGTGGCCCTGCTCAGTGGCA ATCAGCTCCACTGCGGAGGCGTCCTGGTCAATGAGCGCTGGGTGCTCACTGCCGCCCACTGCAAGAT GAATGAGTACACCGTGCACCTGGGCAGTGATACGCTGGGCGACAGGAGAGCTCAGAGGATCAAGGCC TCGAAGTCATTCCGCCACCCCGGCTACTCCACACAGACCCATGTTAATGACCTCATGCTCGTGAAGC TCAATAGCCAGGCCAGGCTGTCATCCATGGTGAAGAAAGTCAGGCTGCCCTCCCGCTGCGAACCCCC TGGAACCACCTGTACTGTCTCCGGCTGGGGCACTACCACGAGCCCAGATGTGACCTTTCCCTCTGAC CTCATGTGCGTGGATGTCAAGCTCATCTCCCCCCAGGACTGCACGAAGGTTTACAAGGACTTACTGG AAAAATTCCATGCTGTGCGCTGGCATCCCCGACTCCAAGAAAAACGCCTGCAATGGTGACTCAGGGGG ACCGTTGGTGTGCAGAGGTACCCTGCAAGGTCTGGTGTCCTGGGGAACTTTCCCTTGCGGCCAACCC AATGACCCAGGAGTCTACACTCAAGTGTGCAAGTTCACCAAGTGGATAAATGACACCATGAAAAAGC ATCGC**TAA**CGCCACACTGAGTTAATTAACTGTGTGCTTCCAACAGAAAATGCACAGGA
	ORF Start: ATG at 54		ORF Stop: TAA at 810

	SEQ ID NO: 94	252 aa	MW at 27557.6kD
NOV18b, CG145754-03 Protein Sequence	MGRPRPRAAKTWMFLLLLGGAWAARGDKIIDGAPCARGSHPWQVALLSGNQLHCGGVLVNERWVLTA AHCKMNEYTVHLGSDTLGDRRAQRIKASKSFRHPGYSTQTHVNDLMLVKLNSQARLSSMVKKVRLPS RCEPPGTTCTVSGWGTTTSPDVTFPSDLMCVDVKLISPQDCTKVYKDLLENSMLCAGIPDSKKNACN GDSGGPLVCRGTLQGLVSWGTFPCGQPNDPGVYTQVCKFTKWINDTMKKHR

	SEQ ID NO: 95	804 bp	
NOV18c, CG145754-02 DNA Sequence	GGATTTCCGGGCTCC**ATG**GCAAGATCCCTTCTCCTGCCCCTGCAGATCCTACTGCTATCCTTAGCCT TGGAAACTGCAGGAGAAGAAGCCCAGGGTGACAAGATTATTGATGGCGCCCCATGTGCAAGAGGCTC CCACCCATGGCAGGTGGCCCTGCTCAGTGGCAATCAGCTCCACTGCGGAGGCGTCCTGGTCAATGAG CGCTGGGTGCTCACTGCCGCCCACTGCAAGATGAATGAGTACACCGTGCACCTGGGCAGTGATACGC TGGGCGACAGGAGAGCTCAGAGGATCAAGGCCTCGAAGTCATTCCGCCACCCCGGCTACTCCACACA GACCCATGTTAATGACCTCAAGCTCATCTCCCCCCAGGACTGCACGAAGGTTTACAAGGACTTACTG GAAAAATTCCATGCTGTGCGCTGGCATCCCCGACTCCAAGAAAAACGCCTGCAATGGTGACTCAGGGG GACCGTTGGTGTGCAGAGGTACCCTGCAAGGTCTGGTGTCCTGGGGAACTTTCCCTTGCGGCCAACC CAATGACCCAGGAGTCTACACTCAAGTGTGCAAGTTCACCAAGTGGATAAATGACACCATGAAAAAG CATCGC**TAA**CGCCACACTGAGTTAATTAACTGTGTGCTTCCAACAGAAAATGCACAGGAGTGAGGAC GCCGATGACCTATGAAGTCAAATTTGACTTTACCTTTCCTCAAAGATATATTTAAACCTCATGCCCT GTTGATAAACCAATCAAATTGGTAAAGACCTAAAACCAAAACAAATAAAGAAACACAAAACCCTCAA
	ORF Start: ATG at 16		ORF Stop: TAA at 610

	SEQ ID NO: 96	198 aa	MW at 21613.6kD
NOV18c, CG145754-02 Protein Sequence	MARSLLLPLQILLLSLALETAGEEAQGDKIIDGAPCARGSHPWQVALLSGNQLHCGGVLVNERWVLT AAHCKMNEYTVHLGSDTLGDRRAQRIKASKSFRHPGYSTQTHVNDLKLISPQDCTKVYKDLLENSML CAGIPDSKKNACNGDSGGPLVCRGTLQGLVSWGTFPCGQPNDPGVYTQVCKFTKWINDTMKKHR

	SEQ ID NO: 97	544 bp
NOV18d, 252718128 DNA Sequence	CACCGGATCCGAAGAAGCCCAGGGTGACAAGATTATTGATGGCGCCCCATGTGCAAGAGGCTCCCAC CCATGGCAGGTGGCCCTGCTCAGTGGCAATCAGCTCCACTGCGGAGGCGTCCTGGTCAATGAGCGCT GGGTGCTCACTGCCGCCCACTGCAAGATGAATGAGTACACCGTGCACCTGGGCAGTGATACGCTGGG CGACAGGAGAGCTCAGAGGATCAAGGCCTCGAAGTCATTCCGCCACCCCGGCTACTCCACACAGACC CATGTTAATGACCTCAAGCTCATCTCCCCCCAGGACTGCACGAAGGTTTACAAGGACTTACTGGAAA ATTCCATGCTGTGCGCTGGCATCCCCGACTCCAAGAAAAACGCCTGCAATGGTGACTCAGGGGGACC GTTGGTGTGCAGAGGTACCCTGCAAGGTCTGGTGTCCTGGGGAACTTTCCCTTGCGGCCAACCCAAT GACCCAGGAGTCTACACTCAAGTGTGCAAGTTCACCAAGTGGATAAATGACACCATGAAAAAGCATC TCGAGGGC	
	ORF Start: at 2	ORF Stop: end of sequence

5

	SEQ ID NO: 98	181 aa	MW at 19683.2kD
NOV18d, 252718128 Protein Sequence	TGSEEAQGDKIIDGAPCARGSHPWQVALLSGNQLHCGGVLVNERWVLTAAHCKMNEYTVHLGSDTLG DRRAQRIKASKSFRHPGYSTQTHVNDLKLISPQDCTKVYKDLLENSMLCAGIPDSKKNACNGDSGGP LVCRGTLQGLVSWGTFPCGQPNDPGVYTQVCKFTKWINDTMKKHLEG		

10

	SEQ ID NO: 99	292 bp
NOV18e, 252718152 DNA Sequence	CACCGGATCCGAAGAAGCCCAGGGTGACAAGATTATTGATGGCGCCCCATGTGCAAGAGGCTCCCAC CCATGGCAGGTGGCCCTGCTCAGTGGCAATCAGCTCCACTGCGGAGGCGTCCTGGTCAATGAGCGCT GGGTGCTCACTGCCGCCCACTGCAAGATGAATGAGTACACCGTGCACCTGGGCAGTGATACGCTGGG CGACAGGAGAGCTCAGAGGATCAAGGCCTCGAAGTCATTCCGCCACCCCGGCTACTCCACACAGACC CATGTTAATGACCTCCTCGAGGGC	
	ORF Start: at 2	ORF Stop: end of sequence

15

	SEQ ID NO: 100	97 aa	MW at 10551.7kD
NOV18e, 252718152 Protein Sequence	TGSEEAQGDKIIDGAPCARGSHPWQVALLSGNQLHCGGVLVNERWVLTAAHCKMNEYTVHLGSDTLG DRRAQRIKASKSFRHPGYSTQTHVNDLLEG		

20

	SEQ ID NO: 101	742 bp
NOV18f, 247856668 DNA Sequence	AGGCTCCGCGGCCGCCCCCTTCACCGGATCCGCCAGGGGTGACAAGATTATTGATGGCGCCCCATGT GCAAGAGGCTCCCACCCATGGCAGGTGGCCCTGCTCAGTGGCAATCAGCTCCACTGCGGAGGCGTCC TGGTCAATGAGCGCTGGGTGCTCACTGCCGCCCACTGCAAGATGAATGAGTACACCGTGCACCTGGG CAGTGATACGCTGGGCGACAGGAGAGCTCAGAGGATCAAGGCCTCGAAGTCATTCCGCCACCCCGGC TACTCCACACAGACCCATGTTAATGACCTCATGCTCGTGAAGCTCAATAGCCAGGCCAGGCTGTCAT CCATGGTGAAGAAAGTCAGGCTGCCCTCCCGCTGCGAACCCCCTGGAACCACCTGTACTGTCTCCGG CTGGGGCACTACCACGAGCCCAGATGTGACCTTTCCCTCTGACCTCATGTGCGTGGATGTCAAGCTC	

	ATCTCCCCCCAGGACTGCACGAAGGTTTACAAGGACTTAGTGGAAAATTCCATGCTGTGCGCTGGCA TCCCCGACTCCAAGAAAAACGCCTGCAATGGTGACTCAGGGGGACCGTTGGTGTGCAGAGGTACCCT GCAAGGTCTGGTGTCCTGGGGAACTTTCCCTTGCGGCCAACCCAATGACCCAGGAGTCTACACTCAA GTGTGCAAGTTCACCAAGTGGATAAATGACACCATGAAAAAGCATCGCCTCGAGGGCAAGGGTGGGC GCGCC
	ORF Start: at 2	ORF Stop: end of sequence

	SEQ ID NO: 102	247 aa	MW at 26591.2kD
NOV18f, 247856668 Protein Sequence	GSAAAPFTGSARGDKIIDGAPCARGSHPWQVALLSGNQLHCGGVLVNERWVLTAAHCKMNEYTVHLG SDTLGDRRAQRIKASKSFRHPGYSTQTHVNDLMLVKLNSQARLSSMVKKVRLPSRCEPPGTTCTVSG WGTTTSPDVTFPSDLMCVDVKLISPQDCTKVYKDLLENSMLCAGIPDSKKNACNGDSGGPLVCRGTL QGLVSWGTFPCGQPNDPGVYTQVCKFTKWINDTMKKHRLEGKGGRA		

	SEQ ID NO: 103	673 bp
NOV18g, 247856705 DNA Sequence	AGGCTCCGCGGCCGCCCCCTTCACCGGATCCGCCAGGGGTGACAAGATTATTGATGGCGCCCCATGT GCAAGAGGCTCCCACCCATGGCAGGTGGCCCTGCTCAGTGGCAATCAGCTCCACTGCGGAGGCGTCC TGGTCAATGAGCGCTGGGTGCTCACTGCCGCCCACTGCAAGATGAATGAGTACACCGTGCACCTGGG CAGTGATACGCTGGGCGACAGGAGAGCTCAGAGGATCAAGGCCTCGAAGTCATTCCGCCACCCCGGC TACTCCACACAGACCCATGTTAATGACCTCATGCTCGTGAAGCTCAATAGCCAGGCCAGGCTGTCAT CCATGGTGAAGAAAGTCAGGCTGCCCTCCCGCTGCGAACCCCCTGGAACCACCTGTACTGTCTCCGG CTGGGGCACTACCACGAGCCCAGATGTGACCTTTCCCTCTGACCTCATGTGCGTGGATGTCAAGCTC ATCTCCCCCCAGGACTGCACGAAGGTTTACAAGGACTTACTGGAAAATTCCATGCTGTGCGCTGGCA TCCCCGACTCCAAGAAAAACGCCTGCAATGGTGACTCAGGGGGACCGTTGGTGTGCAGAGGTACCCT GCAAGGTCTGGTGTCCTGGGGAACTTTCCCTTGCGGCCAACCCAATCTCGAGGGCAAGGGTGGGCGC GCC
	ORF Start: at 2	ORF Stop: end of sequence

	SEQ ID NO: 104	224 aa	MW at 23813.0kD
NOV18g, 247856705 Protein Sequence	GSAAAPFTGSARGDKIIDGAPCARGSHPWQVALLSGNQLHCGGVLVNERWVLTAAHCKMNEYTVHLG SDTLGDRRAQRIKASKSFRHPGYSTQTHVNDLMLVKLNSQARLSSMVKKVRLPSRCEPPGTTCTVSG WGTTTSPDVTFPSDLMCVDVKLISPQDCTKVYKDLLENSMLCAGIPDSKKNACNGDSGGPLVCRGTL QGLVSWGTFPCGQPNLEGKGGRA		

Sequence comparison of the above protein sequences yields the following sequence relationships shown in Table 18B.

20

Table 18B. Comparison of NOV18a against NOV18b through NOV18g.		
Protein Sequence	**NOV18a Residues/ Match Residues**	**Identities/ Similarities for the Matched Region**
NOV18b	25..250 27..252	213/226 (94%) 213/226 (94%)

NOV18c	16..250 19..198	176/235 (74%) 177/235 (74%)
NOV18d	17..249 1..178	172/233 (73%) 173/233 (73%)
NOV18e	17..111 1..95	92/95 (96%) 93/95 (97%)
NOV18f	22..250 11..239	215/229 (93%) 216/229 (93%)
NOV18g	22..230 11..219	193/209 (92%) 194/209 (92%)

Further analysis of the NOV18a protein yielded the following properties shown in Table 18C.

5

Table 18C. Protein Sequence Properties NOV18a	
PSort analysis:	0.6233 probability located in outside; 0.1000 probability located in endoplasmic reticulum (membrane); 0.1000 probability located in endoplasmic reticulum (lumen); 0.1000 probability located in microbody (peroxisome)
SignalP analysis:	Cleavage site between residues 20 and 21

A search of the NOV18a protein against the Geneseq database, a proprietary
10   database that contains sequences published in patents and patent publication, yielded several homologous proteins shown in Table 18D.

Table 18D. Geneseq Results for NOV18a				
Geneseq Identifier	Protein/Organism/Length [Patent #, Date]	NOV18a Residues/ Match Residues	Identities/ Similarities for the Matched Region	Expect Value
AAU82740	Amino acid sequence of novel human protease #39 - Homo sapiens, 253 aa. [WO200200860-A2, 03-JAN-2002]	1..250 4..253	250/250 (100%) 250/250 (100%)	e-150

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AAW05383	Human amyloid precursor protein protease - Homo sapiens, 253 aa. [WO9631122-A1, 10-OCT-1996]	1..250 4..253	250/250 (100%) 250/250 (100%)	e-150
AAR67888	Human stratum corneum chymotrophic recombinant enzyme (SCCE) - Homo sapiens, 253 aa. [WO9500651-A, 05-JAN-1995]	1..250 4..253	250/250 (100%) 250/250 (100%)	e-150
AAB21326	Human HSCEE - Homo sapiens, 257 aa. [WO200053776-A2, 14-SEP-2000]	1..250 4..257	249/255 (97%) 249/255 (97%)	e-146
AAB98502	Human Stratum Corneum Chymotryptic Enzyme, SCCE, catalytic domain - Homo sapiens, 225 aa. [WO200129056-A1, 26-APR-2001]	26..250 1..225	225/225 (100%) 225/225 (100%)	e-136

In a BLAST search of public sequence datbases, the NOV18a protein was found to have homology to the proteins shown in the BLASTP data in Table 18E.

5

Table 18E. Public BLASTP Results for NOV18a				
Protein Accession Number	Protein/Organism/Length	NOV18a Residues/ Match Residues	Identities/ Similarities for the Matched Portion	Expect Value
P49862	Kallikrein 7 precursor (EC 3.4.21.-) (Stratum corneum chymotryptic enzyme) (hSCCE) - Homo sapiens (Human), 253 aa.	1..250 4..253	250/250 (100%) 250/250 (100%)	e-149
AAH32005	Kallikrein 7 (chymotryptic, stratum corneum) - Homo sapiens (Human), 253 aa.	1..250 4..253	249/250 (99%) 249/250 (99%)	e-148
Q91VE3	Thymopsin (Stratum corneum chymotryptic enzyme) - Mus musculus (Mouse), 249 aa.	3..250 5..249	185/248 (74%) 212/248 (84%)	e-111

| AAN03663 | Kallikrein 7 short variant protein - Homo sapiens (Human), 181 aa. | 70..250 1..181 | 181/181 (100%) 181/181 (100%) | e-107 |
| Q9R048 | Stratum corneum chymotryptic enzyme - Mus musculus (Mouse), 234 aa (fragment). | 3..235 5..234 | 175/233 (75%) 198/233 (84%) | e-102 |

PFam analysis predicts that the NOV18a protein contains the domains shown in the Table 18F.

5

Table 18F. Domain Analysis of NOV18a			
Pfam Domain	NOV18a Match Region	Identities/ Similarities for the Matched Region	Expect Value
trypsin	27..242	93/262 (35%) 182/262 (69%)	3.8e-87

Example 19.

10   The NOV19 clone was analyzed, and the nucleotide and encoded polypeptide sequences are shown in Table 19A.

Table 19A. NOV19 Sequence Analysis			
	SEQ ID NO: 105	2028 bp	
NOV19a, CG146279-01 DNA Sequence	TTGAGGACTTTATTATTATTTGGGTTCTTTTCATTTCTTCCCCTTCTGGGCAACGAAGCAATGAAAT TTCCAATCGAGACGCCAAGAAAACAGGTGAACTGGGATCCTAAAGTGGCCGTTCCCGCAGCAGCACC GGTGTGCCAGCCCAAGAGCGCCACTAACGGGCAACCCCCGGCTCCGGCTCCGACTCCAACTCCGCGC CTGTCCATTTCCTCCCGAGCCACAGTGGTAGCCAGGATGGAAGGCACCTCCCAAGGGGGCTTGCAGA CCGTCATGAAGTGGAAGACGGTGGTTGCCATCTTTGTGGTTGTGGTGGTCTACCTTGTCACTGGCGG TCTTGTCTTCCGGGGCATTGGAGCAGCCCTTTGAGAGCAGCCAGAAGAATACCATCGCCTTGGAGAAG GCGGAATTCCTGCGGGATCATGTCTGTGTGAGCCCCCAGGAGCTGGAGACGTTGATCCAGCATGCTC TTGATGCTGACAATGCGGGAGTCAGTCCAATAGGAAACTCTTCCAACAACAGCAGCCACTGGGACCT CGGCAGTGCCTTTTTTCTTTGCTGGAACTGTCATTACGACCATAGGGTATGGGAATATTGCTCCGAGC ACTGAAGGAGGCAAAATCTTTTGTATTTTATATGCCATCTTTGGAATTCCACTCTTTGGTTTCTTAT TGGCTGGAATTGGAGACCAACTTGGAACCATCTTTGGGAAAAGCATTGCAAGAGTGGAGAAGGTCTT TCGAAAAAAGCAAGTGAGTCAGACCAAGATCCGGGTCATCTCAACCATCCTGTTCATCTTGGCCGGC TGCATTGTGTTTGTGACGATCCCTGCTGTCATCTTTAAGTACATCGAGGGCTGGACGGCCTTGGAGT CCATTTACTTTGTGGTGGTCACTCTGACCACGGTGGGCTTTGGTGATTTTGTGGCAGGGGGAAACGC TGGCATCAATTATCGGGAGTGGTATAAGCCCCTAGTGTGGTTTTGGATCCTTGTTGGCCTTGCCTAC TTTGCAGCTGTCCTCAGTATGATCGGAGATTGGCTACGGGTTCTGTCCAAAAAGACAAAAGAAGAGG TGGGTGAAATCAAGGCCCATGCGGCAGAGTGGAAGGCCAATGTCACGGCTGAGTTCCGGGAGACACG GCGAAGGCTCAGCGTGGAGATCCACGATAAGCTGCAGCGGGCGGCCACCATCCGCAGCATGGAGCGC CGGCGGCTGGGCCTGGACCAGCGGGCCCACTCACTGGACATGCTGTCCCCCGAGAAGCGCTCTGTCT TTGCTGCCCTGGACACCGGCCGCTTCAAGGCCTCATCCCAGGAGAGCATCAACAACCGGCCCAACAA CCTGCGCCTGAAGGGGCCGGAGCAGCTGAACAAGCATGGGCAGGGTGCGTCCGAGGACAACATCATC AACAAGTTCGGGTCCACCTCCAGACTCACCAAGAGGAAAAACAAGGACCTCAAAAAGACCTTGCCCG AGGACGTTCAGAAAATCTACAAGACCTTCCGGAATTACTCCCTGGACGAGGAGAAGAAAGAGGAGGA GACGGAAAAGATGTGTAACTCAGACAACTCCAGCACAGCCATGCTGACGGACTGTATCCAGCAGCAC

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| | GCTGAGTTGGAGAACGGAATGATACCCACGGACACCAAAGACCGGGAGCCGGAGAACAACTCATTAC
TTGAAGACAGAAACTAAA<u>ATGTGAAGGACATTGGTCTTGGACTGAGCGTTGTGTGTGTGTGTGTGT</u>
<u>GTTTTTAATATTCACACTGAGACATGTGCCTTAAACAGACTTTTTAGTCCAAAATTACATAGCATTG</u>
<u>AAGAATATATTTCACTGTGCCATAAACAACTGAAAGCTTGCTCTGCCAAAAGGAATCAGAGAACAAG</u>
<u>AACTTCATTTCAGATAGCAAACGCAGGACACACCAAGAGTGTCCGTGCACGTAGCCGGTTCTGGCCG</u>
<u>TACATGTTAAGGGCATTTCAGTGGCAGTGCTGTACCCCTGGGCAGTGCTACCTGGGCACACACGTAG</u>
<u>ACAAGGGCAGCTATTCCT</u> |
| ORF Start: ATG at 61 | | ORF Stop: TAA at 1690 |

	SEQ ID NO: 106	543 aa	MW at 60334.6kD
NOV19a, CG146279-01 Protein Sequence	MKFPIETPRKQVNWDPKVAVPAAAPVCQPKSATNGQPPAPAPTPTPRLSISSRATVVARMEGTSQGG LQTVMKWKTVVAIFVVVVVYLVTGGLVFRALEQPFESSQKNTIALEKAEFLRDHVCVSPQELETLIQ HALDADNAGVSPIGNSSNNSSHWDLGSAFFFAGTVITTIGYGNIAPSTEGGKIFCILYAIFGIPLFG FLLAGIGDQLGTIFGKSIARVEKVFRKKQVSQTKIRVISTILFILAGCIVFVTIPAVIFKYIEGWTA LESIYFVVVTLTTVGFGDFVAGGNAGINYREWYKPLVWFWILVGLAYFAAVLSMIGDWLRVLSKKTK EEVGEIKAHAAEWKANVTAEFRETRRRLSVEIHDKLQRAATIRSMERRRLGLDQRAHSLDMLSPEKR SVFAALDTGRFKASSQESINNRPNNLRLKGPEQLNKHGQGASEDNIINKFGSTSRLTKRKNKDLKKT LPEDVQKIYKTFRNYSLDEEKKEEETEKMCNSDNSSTAMLTDCIQQHAELENGMIPTDTKDREPENN SLLEDRN

Further analysis of the NOV19a protein yielded the following properties shown in Table 19B.

10

Table 19B. Protein Sequence Properties NOV19a	
PSort analysis:	0.6000 probability located in plasma membrane; 0.4000 probability located in Golgi body; 0.3000 probability located in endoplasmic reticulum (membrane); 0.3000 probability located in microbody (peroxisome)
SignalP analysis:	No Known Signal Sequence Predicted

A search of the NOV19a protein against the Geneseq database, a proprietary

15 database that contains sequences published in patents and patent publication, yielded

several homologous proteins shown in Table 19C.

Table 19C. Geneseq Results for NOV19a				
Geneseq Identifier	Protein/Organism/Length [Patent #, Date]	NOV19a Residues/ Match Residues	Identities/ Similarities for the Matched Region	Expect Value

AAU81354	Novel human ion channel protein #34 - Homo sapiens, 543 aa. [WO200185788-A2, 15-NOV-2001]	1..543 1..543	543/543 (100%) 543/543 (100%)	0.0
AAU79472	Human novel transporter protein - Homo sapiens, 543 aa. [WO200224748-A2, 28-MAR-2002]	1..543 1..543	543/543 (100%) 543/543 (100%)	0.0
AAU79473	Human novel transporter protein variant - Homo sapiens, 543 aa. [WO200224748-A2, 28-MAR-2002]	1..543 1..543	542/543 (99%) 543/543 (99%)	0.0
AAE16596	Human TWIK-Related K+ channel-2 (TREK-2) protein - Homo sapiens, 538 aa. [WO200200715-A2, 03-JAN-2002]	18..543 13..538	526/526 (100%) 526/526 (100%)	0.0
AAB47930	Human TREK2 - Homo sapiens, 538 aa. [WO200200715-A2, 03-JAN-2002]	18..543 13..538	526/526 (100%) 526/526 (100%)	0.0

In a BLAST search of public sequence datbases, the NOV19a protein was found to have homology to the proteins shown in the BLASTP data in Table 19D.

5

Table 19D. Public BLASTP Results for NOV19a				
Protein Accession Number	Protein/Organism/Length	NOV19a Residues/ Match Residues	Identities/ Similarities for the Matched Portion	Expect Value
Q8TDK7	Potassium channel TREK2 splice variant b - Homo sapiens (Human), 543 aa.	1..543 1..543	542/543 (99%) 542/543 (99%)	0.0
P57789	Potassium channel subfamily K member 10 (Outward rectifying potassium channel protein TREK-2) (TREK-2 K+ channel subunit) - Homo sapiens (Human), 538 aa.	18..543 13..538	526/526 (100%) 526/526 (100%)	0.0
Q8TDK8	Potassium channel TREK2 splice variant a - Homo sapiens (Human), 543 aa.	18..543 18..543	525/526 (99%) 525/526 (99%)	0.0

| Q9JIS4 | Potassium channel subfamily K member 10 (Outward rectifying potassium channel protein TREK-2) (TREK-2 K+ channel subunit) - Rattus norvegicus (Rat), 538 aa. | 1..543
1..538 | 520/544 (95%)
529/544 (96%) | 0.0 |
| P97438 | Potassium channel subfamily K member 2 (Outward rectifying potassium channel protein TREK-1) (Two-pore potassium channel TPKC1) (TREK-1 K+ channel subunit) - Mus musculus (Mouse), 411 aa. | 22..404
2..369 | 247/384 (64%)
301/384 (78%) | e-136 |

PFam analysis predicts that the NOV19a protein contains the domains shown in the Table 19E.

5

Table 19E. Domain Analysis of NOV19a			
Pfam Domain	**NOV19a Match Region**	**Identities/ Similarities for the Matched Region**	**Expect Value**
ion_trans	158..323	41/231 (18%) 119/231 (52%)	0.046

Example 20.

10   The NOV20 clone was analyzed, and the nucleotide and encoded polypeptide

sequences are shown in Table 20A.

Table 20A. NOV20 Sequence Analysis			
	SEQ ID NO: 107	2958 bp	
NOV20a, CG146374-01 DNA Sequence	GCTCCTCCCCGCTGGCGGGGGGAGAAAGGGCAGGAGGCCTTCCGTCCCGGCTATAAAGGGCCCCGGA CCGCCGCGGCTCGCCTCGGCTTGCCTCGACACGCCTAGGCGCCCTCCGGCTCCGCCCTAGCCGCCGC GTCCCAGCTAGAGCTCCAGCGCCCGCTCAGGCCCCACTCGACCCTCTCGGGCCTCGGCTACTTGGAC TGCGGCGGAATATGGCGGCTCCGATGACTCCCGCGGCTCGGCCCGAGGACTACGAGGCGGCGCTCAA TGCCGCCCTGGCTGACGTGCCCGAACTGGCCAGACTCCTGGAGATCGACCCGTACTTGAAGCCCTAC GCCGTGGACTTCCAGCGCAGGTATAAGCAGTTTAGCCAAATTTTGAAGAACATTGGAGAAAATGAAG GTGGTATTGATAAGTTTTCCAGAGGCTATGAATCATTTGGCGTCCACAGATGTGCTGATGGTGGTTT ATACTGCAAAGAATGGGCCCCGGGAGCAGAAGGAGTTTTTCTTACTGGAGATTTTAATGGTTGGAAT CCATTTTCGTACCCATACAAAAAACTGGATTATGAAAATGGGAGCTGTATATCCCACCAAAGCAGA ATAAATCTGTACTCGTGCCTCATGGATCCAAATTAAAGGTAGTTATTACTAGTAAAGCGGAGAGAT CTTGTATCGTATTTCACCGTGGGCAAAGTATGTGGTTCGTGAAGGTGATAATGTGAATTATGATTGG ATACACTGGGATCCAGAACACTCATATGAGTTTAAGCATTCCAGACCAAAGAAGCCACGGAGTCTAA GAATTTATGAATCTCATGTGGGAATTTCTTCCCATGAAGGAAAAGTAGCTTCTTATAAACATTTTAC ATGCAATGTACTACCAAGAATCAAAGGCCTTGGATACAACTGCATTCAGTTGATGGCAATCATGGAG

CATGCTTACTATGCCAGCTTTGGTTACCAAATCACAAGGTTGTTTGCAGCTTCCAGCCGTTATCCAT
CACCTGAAGAGCTACAAGAACTGGTAGACACAGCTCATTCCATGGGTATCATAGTCCTCTTAGATGT
GGTACACAGCCATGCTTCAAAAAATTCAGCAGATGGATTGAATATGTTTGATGGGACAGATTCCTGT
TATTTTCATTCTGGACCTAGAGGGACTCATGATCTTTGGGATAGCAGATTGTTTGCCTACTCCAGGT
TGAATATTTCAGACATC**TAA**GCCAATTAGAATCATGATTGTTTTGATTGCCAGAAATCCTTAAATCT
GGGAAGTTTTAAGATTCCTTCTGTCAAACATAAGATGGTGGTTGGAAGAATATCGCTTTGATGGATT
TCGTTTTGATGGTGTTACGTCCATGCTTTATCATCACCATGGAGTGGGTCAAGGTTTCTCAGGTGAT
TACAGTGAATATTTCGGACTACAAGTAGATGAAGATGCCTTGACTTACCTCATGTTGGCAAATCATT
TGGTTCACACGCTGTGTCCCGATTCTATAACAATAGCTGAGGATGTATCAGGAATGCCAGCTCTGTG
CTCTCCAATTTCCCAGGGAGGGGGTGGTTTTGACTATCGACTAGCCATGGCAATTCCAGATAAGTGG
ATTCAGCTACTTAAAGAGTTTAAAGATGAAGACTGGAACATGGGCGATATAGTATACACGCTCACAA
ACAGGCGCTACCTTGAAAAGTGCATTGCTTATGCAGGAGCCATGATCAGGCATTGGTTGGGGATAA
GTCGCTGGCATTTTGGTTGATGGATGCCGAAATGTATACAAACATGAGTGTCCTGACTCCTTTTACT
CCAGTTATTGATCGTGGAATACAGCTTCATAAAATGATTCGACTCATTACGCATGGGCTTGGTGGAG
AAGGCTATCTCAATTTCATGGGTAATGAATTTGGGCATCCTGAATGGTTAGACTTCCCAAGAAAAGG
AAATAATGAGAGTTACCATTATGCCAGGCGGCAGTTTCATTTAACTGACGACGACCTTCTTCGCTAC
AAGTTCCTAAATAATTTTGACAGGGATATGAATAGATTGGAAGAAAGATATGGTTGGCTTGCAGCTC
CACAGGCCTACGTGAGTGAAAAACATGAAGGCAATAAGATCATTGCTTTTGAAAGAGCAGGTCTTCT
TTTCATTTTCAACTTCCATCCAAGCAAGAGCTACACTGACTACCGAGTTGGAACAGCATTGCCAGGG
AAATTCAAAATTGTGCTAGATTCAGATGCAGCGGAATATGGAGGGCATCAGAGACTGGACCACAGCA
CTGACTTTTTTTTCTGAGGCTTTTGAACATAATGGGCGTCCCTATTCTCTTTTGGTGTACATTCCAAG
CAGAGTGGCCCTCATCCTTCAGAATGTGGATCTGCCGAATTGAAGAGGCCTGATTTCAGCTCCACCA
GATGCAGATTTGTGTTTTGTTTTCTTGTTATCACTGTCACACAGCTTATAACATGTATGCTTTTCAG
AATACAGTTGTCTAGCCAAGCCATCAAGTGTCTGAAATTCAATATTGGTTTATGCAAATACAGCAAA
CTTTTATTTAAGTAGATAGGAGAATATGTTTAAAATATTAGGAATCCTAGACCATATTTTCAAGTCA
TCTTAGCAGCTAGGATTCTCAAATGGAAGTGGTTATATATAATATGTTAAAAACATTTTGCTTTCCTG
GCTAATTATTTGATCCTTTTAAATTCAAATTTGAATCATTTGTCATGTATGATTATTTCTGTTAAAT
GTACACAGTATTTAAGATGGATATTTGGTGGCTCTATTTGTTCTGATATCTTTTGGTCTAAATTATG
AGGTACCAAGATTGTTTCTTTGTTTCTTTTTTTTTCAAATTGTGTTTAGAAATACTGTAATAAATATGC
AGTAGTGATATAAAGAATTATATCCAAGGTAATATAAAAGCCATTACGTATGAACTCAAAAAAAAAA
AAAAAAAAAA

	ORF Start: ATG at 213		ORF Stop: TAA at 1224

	SEQ ID NO: 108	337 aa	MW at 38247.8kD
NOV20a, CG146374-01 Protein Sequence	MAAPMTPAARPEDYEAALNAALADVPELARLLEIDPYLKPYAVDFQRRYKQFSQILKNIGENEGGID KFSRGYESFGVHRCADGGLYCKEWAPGAEGVFLTGDFNGWNPFSYPYKKLDYGKWELYIPPKQNKSV LVPHGSKLKVVITSKSGEILYRISPWAKYVVREGDNVNYDWIHWDPEHSYEFKHSRPKKPRSLRIYE SHVGISSHEGKVASYKHFTCNVLPRIKGLGYNCIQLMAIMEHAYYASFGYQITSFFAASSRYGSPEE LQELVDTAHSMGIIVLLDVVHSHASKNSADGLNMFDGTDSCYFHSGPRGTHDLWDSRLFAYSRLNIS DI		

Further analysis of the NOV20a protein yielded the following properties shown in Table 20B.

10

Table 20B. Protein Sequence Properties NOV20a	
PSort analysis:	0.7480 probability located in microbody (peroxisome); 0.6000 probability located in nucleus; 0.1000 probability located in mitochondrial matrix space; 0.1000 probability located in lysosome (lumen)
SignalP analysis:	No Known Signal Sequence Predicted

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A search of the NOV20a protein against the Geneseq database, a proprietary database that contains sequences published in patents and patent publication, yielded several homologous proteins shown in Table 20C.

5

Table 20C. Geneseq Results for NOV20a				
Geneseq Identifier	Protein/Organism/Length [Patent #, Date]	NOV20a Residues/ Match Residues	Identities/ Similarities for the Matched Region	Expect Value
AAB90803	Human shear stress-response protein SEQ ID NO: 106 - Homo sapiens, 702 aa. [WO200125427-A1, 12-APR-2001]	1..330 1..330	328/330 (99%) 329/330 (99%)	0.0
ABB60350	Drosophila melanogaster polypeptide SEQ ID NO 7842 - Drosophila melanogaster, 865 aa. [WO200171042-A2, 27-SEP-2001]	22..329 1..314	170/314 (54%) 227/314 (72%)	e-102
AAB49603	Glycogen branching enzyme amino acid sequence - Aspergillus nidulans, 686 aa. [JP2000279180-A, 10-OCT-2000]	31..329 12..314	175/305 (57%) 228/305 (74%)	1e-98
AAG39093	Arabidopsis thaliana protein fragment SEQ ID NO: 48322 - Arabidopsis thaliana, 721 aa. [EP1033405-A2, 06-SEP-2000]	30..329 22..321	161/302 (53%) 214/302 (70%)	3e-92
AAG39092	Arabidopsis thaliana protein fragment SEQ ID NO: 48321 - Arabidopsis thaliana, 858 aa. [EP1033405-A2, 06-SEP-2000]	30..329 159..458	161/302 (53%) 214/302 (70%)	3e-92

In a BLAST search of public sequence datbases, the NOV20a protein was found to have homology to the proteins shown in the BLASTP data in Table 20D.

10

Table 20D. Public BLASTP Results for NOV20a

Protein Accession Number	Protein/Organism/Length	NOV20a Residues/ Match Residues	Identities/ Similarities for the Matched Portion	Expect Value
Q96EN0	Similar to glucan (1,4-alpha-), branching enzyme 1 (glycogen branching enzyme, Andersen disease, glycogen storage disease type IV) - Homo sapiens (Human), 702 aa.	1..330 1..330	330/330 (100%) 330/330 (100%)	0.0
Q04446	1,4-alpha-glucan branching enzyme (EC 2.4.1.18) (Glycogen branching enzyme) (Brancher enzyme) - Homo sapiens (Human), 702 aa.	1..330 1..330	328/330 (99%) 329/330 (99%)	0.0
Q9D6Y9	2310045H19Rik protein (RIKEN cDNA 2310045H19 gene) - Mus musculus (Mouse), 702 aa.	1..330 1..330	291/330 (88%) 310/330 (93%)	e-179
AAF58416	CG4023-PA - Drosophila melanogaster (Fruit fly), 685 aa.	22..329 1..314	170/314 (54%) 227/314 (72%)	e-102
Q9V6K7	CG4023 protein - Drosophila melanogaster (Fruit fly), 865 aa.	22..329 1..314	170/314 (54%) 227/314 (72%)	e-102

PFam analysis predicts that the NOV20a protein contains the domains shown in the Table 20E.

5

Table 20E. Domain Analysis of NOV20a			
Pfam Domain	NOV20a Match Region	Identities/ Similarities for the Matched Region	Expect Value
isoamylase_N	73..168	31/123 (25%) 64/123 (52%)	5.1e-11

Example 21.

The NOV21 clone was analyzed, and the nucleotide and encoded polypeptide

sequences are shown in Table 21A.

5

Table 21A. NOV21 Sequence Analysis			
	SEQ ID NO: 109	885 bp	
NOV21a, CG146403-01 DNA Sequence	TGGATGCTGGCGGTCCTCTACCTGGTCTGGCTCTATTGGGATAGAAACATACCCAGGGCTGGTGGAA GGCGTTCGGAGTGGATAAGGAACCGGGCAATTTGGAGACAACTAAGGGATTATTATCCTGTCAAGCT GGTGAAAACAGCAGAGCTGCCCCCGGATCGGAACTACGTGCTGGGCGCCCACCCTCATGGGATCATG TGTACAGGCTTCCTCTGTAATTTCTCCACCGAGAGCAATGGCTTCTCCCAGCTCTTCCCGGGGCTCC GGCCCTGGTTAGCCGTGCTGGCTGGCCTCTTCTACCTCCCGGTCTATCGCGACTACATCATGTCCTT TGGTCTCTGTCCGGTGAGCCGCCAGAGCCTGGACTTCATCCTGTCCCAGCCCCAGCTCGGGCAGGCC GTGGTCATCATGGTGGGGGGTGCGCACGAGGCCCTGTATTCAGTCCCCGGGGAGCACTGCCTTACGC TCCAGAAGCGCAAAGGCTTCGTGCGCCTGGCGCTGAGGCACGGGGCGTCCCTGGTGCCCGTGTACTC CTTTGGGGAGAATGACATCTTTAGACTTAAGGCTTTTGCCACAGGCTCCTGGCAGCATTGGTGCCAG CTCACCTTCAAGAAGCTCATGGGCTTCTCTCCTTGCATCTTCTGGGGTCGCGGTCTCTTCTCAGCCA CCTCCTGGGGCCTGCTGCCCTTTGCTGTGCCCATCACCACTGTGGGTGAGCCCATCCCCGTCCCCCA GCGCCTCCACCCCACCGAGGAGGAAGTCAATCACTATCACGCCCTCTACATGACGGCCCTGGAGCAG CTCTTCGAGGAGCACAAGGAAAGCTGTGGGGTCCCCGCTTCCACCTGCCTCACCTTCATCTAGGCCT GGCCGCGGCCTTTC		
	ORF Start: ATG at 4		ORF Stop: TAG at 865

	SEQ ID NO: 110	287 aa	MW at 32641.7kD
NOV21a, CG146403-01 Protein Sequence	MLAVLYLVWLYWDRNIPRAGGRRSEWIRNRAIWRQLRDYYPVKLVKTAELPPDRNYVLGAHPHGIMC TGFLCNFSTESNGFSQLFPGLRPWLAVLAGLFYLPVYRDYIMSFGLCPVSRQSLDFILSQPQLGQAV VIMVGGAHEALYSVPGEHCLTLQKRKGFVRLALRHGASLVPVYSFGENDIFRLKAFATGSWQHWCQL TFKKLMGFSPCIFWGRGLFSATSWGLLPFAVPITTVGEPIPVPQRLHPTEEEVNHYHALYMTALEQL FEEHKESCGVPASTCLTFI		

Further analysis of the NOV21a protein yielded the following properties shown in

Table 21B.

15

Table 21B. Protein Sequence Properties NOV21a	
PSort analysis:	0.5500 probability located in endoplasmic reticulum (membrane); 0.3814 probability located in lysosome (lumen); 0.3200 probability located in microbody (peroxisome); 0.1000 probability located in endoplasmic reticulum (lumen)
SignalP analysis:	Cleavage site between residues 22 and 23

A search of the NOV21a protein against the Geneseq database, a proprietary

20   database that contains sequences published in patents and patent publication, yielded

several homologous proteins shown in Table 21C.

Table 21C. Geneseq Results for NOV21a					
Geneseq Identifier	Protein/Organism/Length [Patent #, Date]	NOV21a Residues/ Match Residues	Identities/ Similarities for the Matched Region	Expect Value
AAM80262	Human protein SEQ ID NO 3908 - Homo sapiens, 223 aa. [WO200157190-A2, 09-AUG-2001]	43..237 29..223	195/195 (100%) 195/195 (100%)	e-115
ABB75677	Breast protein-eukaryotic conserved gene 1 (BSTP-ECG1) protein - Homo sapiens, 388 aa. [WO200208260-A2, 31-JAN-2002]	1..284 101..385	158/285 (55%) 218/285 (76%)	1e-97
AAB66170	Protein of the invention #82 - Unidentified, 388 aa. [WO200078961-A1, 28-DEC-2000]	1..284 101..385	158/285 (55%) 218/285 (76%)	1e-97
AAU29191	Human PRO polypeptide sequence #168 - Homo sapiens, 388 aa. [WO200168848-A2, 20-SEP-2001]	1..284 101..385	158/285 (55%) 218/285 (76%)	1e-97
AAY99421	Human PRO1433 (UNQ738) amino acid sequence SEQ ID NO:292 - Homo sapiens, 388 aa. [WO200012708-A2, 09-MAR-2000]	1..284 101..385	158/285 (55%) 218/285 (76%)	1e-97

5   In a BLAST search of public sequence datbases, the NOV21a protein was found to have homology to the proteins shown in the BLASTP data in Table 21D.

Table 21D. Public BLASTP Results for NOV21a					
Protein Accession Number	Protein/Organism/Length	NOV21a Residues/ Match Residues	Identities/ Similarities for the Matched Portion	Expect Value
Q9UDW7	WUGSC:H_DJ0747G18.5 protein - Homo sapiens (Human), 261 aa (fragment).	43..287 16..261	244/246 (99%) 244/246 (99%)	e-145

CAD38961	Hypothetical protein - Homo sapiens (Human), 434 aa (fragment).	1..284 147..431	158/285 (55%) 218/285 (76%)	3e-97
Q96PD7	Diacylglycerol acyltransferase 2 (Hypothetical 43.8 kDa protein) - Homo sapiens (Human), 388 aa.	1..284 101..385	158/285 (55%) 218/285 (76%)	3e-97
Q9BYE5	GS1999full protein - Homo sapiens (Human), 297 aa.	1..284 10..294	158/285 (55%) 218/285 (76%)	3e-97
Q9DCV3	0610010B06Rik protein (Diacylglycerol acyltransferase 2) - Mus musculus (Mouse), 388 aa.	1..284 101..385	159/285 (55%) 217/285 (75%)	8e-97

Example 22.

The NOV22 clone was analyzed, and the nucleotide and encoded polypeptide sequences are shown in Table 22A.

Table 22A. NOV22 Sequence Analysis			
	SEQ ID NO: 111	1135 bp	
NOV22a, CG146513-01 DNA Sequence	CACAGTAAGAGATTATAGCAAAGCATCTATAATCAACTCAGCTTAAGAAGTTTTGACCTTCTGGTTA GGCTTCTTGCCACAACAGAACAGCACCATAACCATGGCTTTCTTCTCCCGACTGAATCTCCAGGAGG GCCTCCAAACCTTCTTTGTTTTGCAATGGATCCCAGTCTATATATTTTTAGGAGCTATTCCCATTCT CCTTATACCCTACTTTCTGTTATTCAGTAAGTTCTGGCCCTTGGCTGTGCTCTCCTTAGCCTGGCTC ACCTATGATTGGAACACCCACAGTCAAGGTGGCAGGCGTTCAGCTTGGGTACGAAACTGGACCCTAT GGAAGTATTTCCGAAATTACTTCCCAGTACAGCTGGTGAAGACTCATGATCTTTCTCCCAAACACAA CTACATCATTGCCAATCACCCCATGGCATTCTCTCTTTTGGTGTCTTCATCAACTTTGCCACTGAG GCCACTGGCATTGCTCGGATTTTCCCATCCATCACTCCCTTTGTAGGGACCTTAGAAAGGATATTTT GGATCCCAATTGTGCGAGAATATGTGATGTCAATGGGTGTGTGCCCTGTGAGTAGCTCAGCCTTGAA GTACTTGCTGACCCAGAAAGGCTCAGGCAATGCCGTGGTTATTGTGGTGGGTGGAGCTGCTGAAGCT CTCTTGTGCCGACCAGGAGCCTCCACTCTCTTCCTCAAGCAGCGTAAAGGTTTTGTGAAGATGGCAC TGCAAACAGGGGCATACCTTGTCCCTTCATATTCCTTTGGTGAGAACGAAGTTTTCAATCAGGAGAC CTTCCCTGAGGGCACGTGGTTAAGGTTGTTCCAAAAAACCTTCCAGGACACATTCAAAAAAATCCTG GGACTAAATTTCTGTACCTTCCATGGCCGGGGCTTCACTCGCGGATCCTGGGGCTTCCTGCCTTTCA ATCGGCCCATTACCACTGTTGGGGAACCCCTTCCAATTCCCAGGATTAAGAGGCCAAACCAGAAGAC AGTAGACAAGTATCACGCACTCTACATCAGTGCCCTGCGCAAGCTCTTTGACCAACACAAAGTTGAA TATGGCCTCCCTGAGACCCAAGAGCTGACAATTACATAACAGGAGCCACATTCCCCATTGATC
	ORF Start: ATG at 101		ORF Stop: TAA at 1109

	SEQ ID NO: 112	336 aa	MW at 38493.6kD
NOV22a, CG146513-01 Protein Sequence	MAFFSRLNLQEGLQTFFVLQWIPVYIFLGAIPILLIPYFLLFSKFWPLAVLSLAWLTYDWNTHSQGG RRSAWVRNWTLWKYFRNYFPVQLVKTHDLSPKHNYIIANHPHGILSFGVFINFATEATGIARIFPSI TPFVGTLERIFWIPIVREYVMSMGVCPVSSSALKYLLTQKGSGNAVVIVVGGAAEALLCRPGASTLF LKQRKGFVKMALQTGAYLVPSYSFGENEVFNQETFPEGTWLRLFQKTFQDTFKKILGLNFCTFHGRG FTRGSWGFLPFNRPITTVGEPLPIPRIKRPNQKTVDKYHALYISALRKLFDQHKVEYGLPETQELTI T

Further analysis of the NOV22a protein yielded the following properties shown in Table 22B.

Table 22B. Protein Sequence Properties NOV22a	
PSort analysis:	0.6850 probability located in plasma membrane; 0.6400 probability located in endoplasmic reticulum (membrane); 0.3880 probability located in microbody (peroxisome); 0.3700 probability located in Golgi body
SignalP analysis:	Cleavage site between residues 65 and 66

A search of the NOV22a protein against the Geneseq database, a proprietary database that contains sequences published in patents and patent publication, yielded several homologous proteins shown in Table 22C.

Table 22C. Geneseq Results for NOV22a				
Geneseq Identifier	Protein/Organism/Length [Patent #, Date]	NOV22a Residues/ Match Residues	Identities/ Similarities for the Matched Region	Expect Value
AAM06866	Human foetal protein, SEQ ID NO: 1074 - Homo sapiens, 225 aa. [WO200155339-A2, 02-AUG-2001]	1..216 1..216	211/216 (97%) 214/216 (98%)	e-124
ABB75677	Breast protein-eukaryotic conserved gene 1 (BSTP-ECG1) protein - Homo sapiens, 388 aa. [WO200208260-A2, 31-JAN-2002]	1..335 56..387	171/337 (50%) 237/337 (69%)	e-101
AAB66170	Protein of the invention #82 - Unidentified, 388 aa. [WO200078961-A1, 28-DEC-2000]	1..335 56..387	171/337 (50%) 237/337 (69%)	e-101
AAU29191	Human PRO polypeptide sequence #168 - Homo sapiens, 388 aa. [WO200168848-A2, 20-SEP-2001]	1..335 56..387	171/337 (50%) 237/337 (69%)	e-101

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AAY99421	Human PRO1433 (UNQ738) amino acid sequence SEQ ID NO:292 - Homo sapiens, 388 aa. [WO200012708-A2, 09-MAR-2000]	1..335 56..387	171/337 (50%) 237/337 (69%)	e-101

In a BLAST search of public sequence datbases, the NOV22a protein was found to have homology to the proteins shown in the BLASTP data in Table 22D.

5

Table 22D. Public BLASTP Results for NOV22a				
Protein Accession Number	Protein/Organism/Length	NOV22a Residues/ Match Residues	Identities/ Similarities for the Matched Portion	Expect Value
Q9DCV3	0610010B06Rik protein (Diacylglycerol acyltransferase 2) - Mus musculus (Mouse), 388 aa.	1..335 56..387	172/337 (51%) 238/337 (70%)	e-101
CAD38961	Hypothetical protein - Homo sapiens (Human), 434 aa (fragment).	1..335 102..433	171/337 (50%) 237/337 (69%)	e-100
Q96PD7	Diacylglycerol acyltransferase 2 (Hypothetical 43.8 kDa protein) - Homo sapiens (Human), 388 aa.	1..335 56..387	171/337 (50%) 237/337 (69%)	e-100
Q8TAB1	BA351K23.5 (Novel protein) - Homo sapiens (Human), 296 aa (fragment).	38..335 1..295	161/299 (53%) 221/299 (73%)	2e-98
Q9BYE5	GS1999full protein - Homo sapiens (Human), 297 aa.	39..335 2..296	161/299 (53%) 217/299 (71%)	4e-96

10   **Example 23.**

The NOV23 clone was analyzed, and the nucleotide and encoded polypeptide sequences are shown in Table 23A.

Table 23A. NOV23 Sequence Analysis			
	SEQ ID NO: 113	1022 bp	

NOV23a, CG146522-01 DNA Sequence	ACTGTTCTGAGATCTTTGCCTCCCTCAGGCTCCCGAGAATCATGGCTCATTCCAAGCAGCCTAGTCA CTTCCAGAGTCTGATGCTTCTGCAGTGGCCTTTGAGCTACCTTGCCATCTTGTTCGTCTACCTGCTG TTTACATCCTTGTGGCCGCTACCAGTGCTTTACTTTGCCTGGTTGTTCCTGGACTGGAAGACCCCAG AGCGAGGTGGCAGGCGTTCGGCCTGGGTAAGGAACTGGTGTGTCTGGACCCACATCAGGGACTATTT CCCCATTATCCTGAAGACAAAGGACCTATCACCTGAGCACAACTACCTCATGGGGGTTCACCCCCAT GGCCTCCTGACCTTTGGCGCCTTCTGCAACTTCTGCACTGAGGCCACAGGCTTCTCGAAGACCTTCC CAGGCATCACTCCTCACTTGGCCACGCTGTCCTGGTTCTTCAAGATCCCCTTTGTTAGGGAGTACCT CATGGCCAAAGGTGTGTGCTCTGTGAGCCAGCCAGCCATCAACTATCTGCTGAGCCATGGCACTGGC AACCTCGTGGGCATTGTAGTGGGAGGTGTGGGTGAGGCCCTGCAAAGTGTGCCCAACACCACCACCC TCATCCTCCAGAAGCGCAAGGGGTTCGTGCGCACAGCCCTCCAGCATGGGGCTCATCTGGTCCCCAC CTTCACTTTTGGGGAAACTGAGGTGTATGATCAGGTGCTGTTCCATAAGGATAGCAGGATGTACAAG TTCCAGAGCTGCTTCCGCCGTATCTTTGGTTTCTACTGTTGTGTGTCTTCTATGGACAAAGCTTCTGTC AAGGCTCCACTGGGCTCCTGCCATACTCCAGGCCTATTGTCACTGTTGGGGAGCCTCTGCCACTGCC CCAAATTGAAAAGCCAAGCCAGGAGATGGTGGACAAATACCATGCACTTTATATGGATGCTCTGCAC AAACTGTTCGACCAGCATAAGACCCACTATGGCTGCTCAGAGACCCAAAAGCTGTTTTTCCTGTGAA TGAAGGTACTGCATGCC
	ORF Start: ATG at 42		ORF Stop: TGA at 1002

	SEQ ID NO: 114	320 aa	MW at 36773.5kD
NOV23a, CG146522-01 Protein Sequence	MAHSKQPSHFQSLMLLQWPLSYLAILFVYLLFTSLWPLPVLYFAWLFLDWKTPERGGRRSAWVRNWC VWTHIRDYFPIILKTKDLSPEHNYLMGVHPHGLLTFGAFCNFCTEATGFSKTFPGITPHLATLSWFF KIPFVREYLMAKGVCSVSQPAINYLLSHGTGNLVGIVVGGVGEALQSVPNTTTLILQKRKGFVRTAL QHGAHLVPTFTFGETEVYDQVLFHKDSRMYKFQSCFRRIFGFYCCVFYGQSFCQGSTGLLPYSRPIV TVGEPLPLPQIEKPSQEMVDKYHALYMDALHKLFDQHKTHYGCSETQKLFFL		

Further analysis of the NOV23a protein yielded the following properties shown in Table 23B.

10

Table 23B. Protein Sequence Properties NOV23a	
PSort analysis:	0.7284 probability located in outside; 0.3880 probability located in microbody (peroxisome); 0.1000 probability located in endoplasmic reticulum (membrane); 0.1000 probability located in endoplasmic reticulum (lumen)
SignalP analysis:	Cleavage site between residues 43 and 44

A search of the NOV23a protein against the Geneseq database, a proprietary

15   database that contains sequences published in patents and patent publication, yielded

several homologous proteins shown in Table 23C.

Table 23C. Geneseq Results for NOV23a				
Geneseq Identifier	Protein/Organism/Length [Patent #, Date]	NOV23a Residues/ Match Residues	Identities/ Similarities for the Matched Region	Expect Value

ABB75677	Breast protein-eukaryotic conserved gene 1 (BSTP-ECG1) protein - Homo sapiens, 388 aa. [WO200208260-A2, 31-JAN-2002]	4..317 62..385	165/324 (50%) 224/324 (68%)	1e-93
AAB66170	Protein of the invention #82 - Unidentified, 388 aa. [WO200078961-A1, 28-DEC-2000]	4..317 62..385	165/324 (50%) 224/324 (68%)	1e-93
AAU29191	Human PRO polypeptide sequence #168 - Homo sapiens, 388 aa. [WO200168848-A2, 20-SEP-2001]	4..317 62..385	165/324 (50%) 224/324 (68%)	1e-93
AAY99421	Human PRO1433 (UNQ738) amino acid sequence SEQ ID NO:292 - Homo sapiens, 388 aa. [WO200012708-A2, 09-MAR-2000]	4..317 62..385	165/324 (50%) 224/324 (68%)	1e-93
AAY94889	Human protein clone HP02485 - Homo sapiens, 334 aa. [WO200005367-A2, 03-FEB-2000]	11..319 16..333	144/318 (45%) 200/318 (62%)	3e-74

In a BLAST search of public sequence datbases, the NOV23a protein was found to have homology to the proteins shown in the BLASTP data in Table 23D.

5

Table 23D. Public BLASTP Results for NOV23a				
Protein Accession Number	Protein/Organism/Length	NOV23a Residues/ Match Residues	Identities/ Similarities for the Matched Portion	Expect Value
Q8TAB1	BA351K23.5 (Novel protein) - Homo sapiens (Human), 296 aa (fragment).	30..317 3..293	163/291 (56%) 214/291 (73%)	5e-96
Q9DCV3	0610010B06Rik protein (Diacylglycerol acyltransferase 2) - Mus musculus (Mouse), 388 aa.	4..317 62..385	166/324 (51%) 225/324 (69%)	2e-93
CAD38961	Hypothetical protein - Homo sapiens (Human), 434 aa (fragment).	4..317 108..431	165/324 (50%) 224/324 (68%)	3e-93

| Q96PD7 | Diacylglycerol acyltransferase 2 (Hypothetical 43.8 kDa protein) - Homo sapiens (Human), 388 aa. | 4..317 62..385 | 165/324 (50%) 224/324 (68%) | 3e-93 |
| Q9BYE5 | GS1999full protein - Homo sapiens (Human), 297 aa. | 28..317 1..294 | 156/294 (53%) 210/294 (71%) | 1e-89 |

Example 24.

The NOV24 clone was analyzed, and the nucleotide and encoded polypeptide sequences are shown in Table 24A.

Table 24A. NOV24 Sequence Analysis			
	SEQ ID NO: 115	1056 bp	
NOV24a, CG146531-01 DNA Sequence	CATTTTCCAAAGGTGTCACAGGAAGAGCATGGCAGAGCTGGGACTGGGAGCCAGGTCACCATGGCTT TCTTCTCCCGACTGAATCTCCAGGAGGGCCTCCAAACCTTCTTTGTTTTGCAATGGATCCCAGTCTA TATATTTTTAGGTTTGTTCGTCTACCTGCTGTTTACATCCTTGTGGCCGCTACCAGTGCTTTACTTT GCCTGGTTGTTCCTGGACTGGAAGACCCCAGAGCGAGGTGGCAGGCGTTCGGCCTGGGTAAGGAACT GGTGTGTCTGGACCCACATCAGGGACTATTTCCCCATTCAGATCCTGAAGACAAAGGACCTATCACC TGAGCACAACTACCTCATGGGGGTTCACCCCCATGGCCTCCTGACCTTTGGCGCCTTCTGCAACTTC TGCACTGAGGCCACAGGCTTCTCGAAGACCTTCCCAGGCATCACTCCTCACTTGGCCACGCTGTCCT GGTTCTTCAAGATCCCCTTTGTTAGGGAGTACCTCATGGCCAAAGGTGTGTGCTCTGTGAGCCAGCC AGCCATCAACTATCTGCTGAGCCATGGCACTGGCAACCTCGTGGGCATTGTAGTGGGAGGTGTGGGT GAGGCCCTGCAAAGTGTGCCCAACACCACCACCCTCATCCTCCAGAAGCGCAAGGGGTTCGTGCGCA CAGCCCTCCAGCATGGGGGCTCATCTGGTCCCCACCTTCACTTTTGGGGAAACTGAGGTGTATGATCA GGTGCTGTTCCATAAGGATAGCAGGATGTACAAGTTCCAGAGCTGCTTCCGCCGTATCTTTGGTTTC TACTGTTGTGTCTTCTATGGACAAAGCTTCTGTCAAGGCTCCACTGGGCTCCTGCCATACTCCAGGC CTATTGTCACTGTTGGGGAGCCTCTGCCACTGCCCCAAATTGAAAAGCCAAGCCAGGAGATGGTGGA CAAATACCATGCACTTTATATGGATGCTCTGCACAAACTGTTCGACCAGCATAAGACCCACTATGGC TGCTCAGAGACCCAAAAGCTGTTTTTCCTGTGAATGAAGGTACTGCATGCC		
	ORF Start: ATG at 61		ORF Stop: TGA at 1036

	SEQ ID NO: 116	325 aa	MW at 37453.3kD
NOV24a, CG146531-01 Protein Sequence	MAFFSRLNLQEGLQTFFVLQWIPVYIFLGLFVYLLFTSLWPLPVLYFAWLFLDWKTPERGGRRSAWV RNWCVWTHIRDYFPIQILKTKDLSPEHNYLMGVHPHGLLTFGAFCNFCTEATGFSKTFPGITPHLAT LSWFFKIPFVREYLMAKGVCSVSQPAINYLLSHGTGNLVGIVVGGVGEALQSVPNTTTLILQKRKGF VRTALQHGAHLVPTFTFGETEVYDQVLFHKDSRMYKFQSCFRRIFGFYCCVFYGQSFCQGSTGLLPY SRPIVTVGEPLPLPQIEKPSQEMVDKYHALYMDALHKLFDQHKTHYGCSETQKLFFL		

Further analysis of the NOV24a protein yielded the following properties shown in Table 24B.

Table 24B. Protein Sequence Properties NOV24a

PSort analysis:	0.8200 probability located in outside; 0.3880 probability located in microbody (peroxisome); 0.1000 probability located in endoplasmic reticulum (membrane); 0.1000 probability located in endoplasmic reticulum (lumen)
SignalP analysis:	Cleavage site between residues 47 and 48

A search of the NOV24a protein against the Geneseq database, a proprietary database that contains sequences published in patents and patent publication, yielded several homologous proteins shown in Table 24C.

Table 24C. Geneseq Results for NOV24a				
Geneseq Identifier	Protein/Organism/Length [Patent #, Date]	NOV24a Residues/ Match Residues	Identities/ Similarities for the Matched Region	Expect Value
ABB75677	Breast protein-eukaryotic conserved gene 1 (BSTP-ECG1) protein - Homo sapiens, 388 aa. [WO200208260-A2, 31-JAN-2002]	1..322 56..385	166/330 (50%) 230/330 (69%)	2e-96
AAB66170	Protein of the invention #82 - Unidentified, 388 aa. [WO200078961-A1, 28-DEC-2000]	1..322 56..385	166/330 (50%) 230/330 (69%)	2e-96
AAU29191	Human PRO polypeptide sequence #168 - Homo sapiens, 388 aa. [WO200168848-A2, 20-SEP-2001]	1..322 56..385	166/330 (50%) 230/330 (69%)	2e-96
AAY99421	Human PRO1433 (UNQ738) amino acid sequence SEQ ID NO:292 - Homo sapiens, 388 aa. [WO200012708-A2, 09-MAR-2000]	1..322 56..385	166/330 (50%) 230/330 (69%)	2e-96
AAY94889	Human protein clone HP02485 - Homo sapiens, 334 aa. [WO200005367-A2, 03-FEB-2000]	13..324 15..333	147/321 (45%) 200/321 (61%)	2e-75

In a BLAST search of public sequence datbases, the NOV24a protein was found to have homology to the proteins shown in the BLASTP data in Table 24D.

Table 24D. Public BLASTP Results for NOV24a				
Protein Accession Number	Protein/Organism/Length	NOV24a Residues/ Match Residues	Identities/ Similarities for the Matched Portion	Expect Value
Q8TAB1	BA351K23.5 (Novel protein) - Homo sapiens (Human), 296 aa (fragment).	34..322 3..293	163/291 (56%) 215/291 (73%)	1e-97
CAD38961	Hypothetical protein - Homo sapiens (Human), 434 aa (fragment).	1..322 102..431	166/330 (50%) 230/330 (69%)	6e-96
Q9DCV3	0610010B06Rik protein (Diacylglycerol acyltransferase 2) - Mus musculus (Mouse), 388 aa.	1..322 56..385	167/330 (50%) 230/330 (69%)	6e-96
Q96PD7	Diacylglycerol acyltransferase 2 (Hypothetical 43.8 kDa protein) - Homo sapiens (Human), 388 aa.	1..322 56..385	166/330 (50%) 230/330 (69%)	6e-96
Q9BYE5	GS1999full protein - Homo sapiens (Human), 297 aa.	32..322 1..294	157/294 (53%) 211/294 (71%)	1e-91

Example 25.

The NOV25 clone was analyzed, and the nucleotide and encoded polypeptide sequences are shown in Table 25A.

Table 25A. NOV25 Sequence Analysis			
	SEQ ID NO: 117	951 bp	
NOV25a, CG147274-01 DNA Sequence	ATGGGGCTTCGGGCAGGCCCCATCCTGCTTCTGCTGCTGTGGCTGCTGCCAGGGGCCCATTGGGATG TGCTGCCTTCAGAATGCGGCCACTCCAAGGAGGCCGGGAGGATTGTGGGAGGCCAAGACACCCAGGA AGGACGCTGGCCGTGGCAGGTTGGCCTGTGGTTGACCTCAGTGGGGCATGTATGTGGGGGCTCCCTC ATCCACCCACGCTGGGTGCTCACAGCCGCCCACTGCTTCCTGAGGTCTGAGGATCCCGGGCTCTACC ATGTTAAAGTCGGAGGGCTGACACCCTCACTTTCAGAGCCCCACTCGGCCTTGGTGGCTGTGAGGAG GCTCCTGGTCCACTCCTCATACCATGGGACCACCACCAGCGGGGACATTGCCCTGATGGAGCTGGAC TCCCCCTTGCAGGCCTCCCAGTTCAGCCCCATCTGCCTCCCAGGACCCCAGACCCCCCTCGCCATTG GGACCGTGTGCTGGGTAAACGGGCTGGGGCCCACATCACATCCAGCCCTGGCGAGTGTCCTTCAGGA GGTGGCTGTGCCCCTCCTGGACTCGAACATGTGTGAGCTGATGTACCACCTAGGAGAGCCCAGCCTG GCTGGCCAGCGCCTCATCCAGGACGACATGCTCTGTGCTGGCTCTGTCCAGGGCAAGAAAGACTCCT GCCAGGGTGACTCCGGGGGGCCGCTGGTCTGCCCCATCAATGATACGTGGATCCAGGCCGGCATTGT GAGCTGGGGATTCGGCTGTGTGCCCGGCCTTTCCGGCCTGGTGTCTACACCCAGGTGCTAAGCTACACA GACTGGATTCAGAGAACCCTGGCTGAATCTCACTCAGGCATGTCTGGGGCCCGCCCAGGTGCCCCAG GATCCCACTCAGGCACCTCCAGATCCCACCCAGTGCTGCTGCTTGAGCTGTTGACCGTATGCTTGCT TGGGTCCCTG**TGA**	
	ORF Start: ATG at 1		ORF Stop: TGA at 949

	SEQ ID NO: 118	316 aa	MW at 33574.2kD
NOV25a, CG147274-01 Protein Sequence	MGLRAGPILLLLLWLLPGAHWDVLPSECGHSKEAGRIVGGQDTQEGRWPWQVGLWLTSVGHVCGGSL IHPRWVLTAAHCFLRSEDPGLYHVKVGGLTPSLSEPHSALVAVRRLLVHSSYHGTTTSGDIALMELD SPLQASQFSPICLPGPQTPLAIGTVCWVNGLGPTSHPALASVLQEVAVPLLDSNMCELMYHLGEPSL AGQRLIQDDMLCAGSVQGKKDSCQGDSGGPLVCPINDTWIQAGIVSWGFGCARPFRPGVYTQVLSYT DWIQRTLAESHSGMSGARPGAPGSHSGTSRSHPVLLLELLTVCLLGSL		

5

Further analysis of the NOV25a protein yielded the following properties shown in Table 25B.

10

Table 25B. Protein Sequence Properties NOV25a	
PSort analysis:	0.9190 probability located in plasma membrane; 0.3000 probability located in lysosome (membrane); 0.1000 probability located in endoplasmic reticulum (membrane); 0.1000 probability located in endoplasmic reticulum (lumen)
SignalP analysis:	Cleavage site between residues 22 and 23

A search of the NOV25a protein against the Geneseq database, a proprietary database that contains sequences published in patents and patent publication, yielded

15   several homologous proteins shown in Table 25C.

Table 25C. Geneseq Results for NOV25a

Geneseq Identifier	Protein/Organism/Length [Patent #, Date]	NOV25a Residues/ Match Residues	Identities/ Similarities for the Matched Region	Expect Value
AAU98887	Human protease PRTS5 - Homo sapiens, 304 aa. [WO200238744-A2, 16-MAY-2002]	1..316 1..304	304/316 (96%) 304/316 (96%)	0.0
AAW77303	Amino acid sequence of SP002LA, a homologue of HELA2 - Homo sapiens, 289 aa. [WO9836054-A1, 20-AUG-1998]	28..316 1..289	285/289 (98%) 285/289 (98%)	e-171
ABG64545	Human albumin fusion protein #1220 - Homo sapiens, 290 aa. [WO200177137-A1, 18-OCT-2001]	5..275 6..276	121/275 (44%) 168/275 (61%)	1e-63
AAB73945	Human protease T - Homo sapiens, 290 aa. [WO200116293-A2, 08-MAR-2001]	5..275 6..276	121/275 (44%) 168/275 (61%)	1e-63
AAE03821	Human gene 4 encoded secreted protein HWHIH10, SEQ ID NO: 67 - Homo sapiens, 290 aa. [WO200136440-A1, 25-MAY-2001]	5..275 6..276	121/275 (44%) 168/275 (61%)	1e-63

In a BLAST search of public sequence datbases, the NOV25a protein was found to

5   have homology to the proteins shown in the BLASTP data in Table 25D.

Table 25D. Public BLASTP Results for NOV25a

Protein Accession Number	Protein/Organism/Length	NOV25a Residues/ Match Residues	Identities/ Similarities for the Matched Portion	Expect Value
Q91XC4	Similar to distal intestinal serine protease - Mus musculus (Mouse), 310 aa.	1..316 1..310	202/317 (63%) 235/317 (73%)	e-114

Q9QYZ9	Distal intestinal serine protease - Mus musculus (Mouse), 310 aa.	1..316 1..310	201/317 (63%) 233/317 (73%)	e-113
Q9BQR3	Marapsin precursor (EC 3.4.21.-) - Homo sapiens (Human), 290 aa.	5..275 6..276	121/275 (44%) 168/275 (61%)	3e-63
Q8R1A6	RIKEN cDNA 2010001P08 gene - Mus musculus (Mouse), 331 aa.	24..305 41..329	142/293 (48%) 174/293 (58%)	5e-62
Q9DGR3	Embryonic serine protease-1 - Xenopus laevis (African clawed frog), 317 aa.	25..304 29..308	123/288 (42%) 165/288 (56%)	1e-59

PFam analysis predicts that the NOV25a protein contains the domains shown in the Table 25E.

Table 25E. Domain Analysis of NOV25a

Pfam Domain	NOV25a Match Region	Identities/ Similarities for the Matched Region	Expect Value
trypsin	37..271	109/266 (41%) 176/266 (66%)	1.7e-79

Example 26.

The NOV26 clone was analyzed, and the nucleotide and encoded polypeptide sequences are shown in Table 26A.

Table 26A. NOV26 Sequence Analysis

	SEQ ID NO: 119	970 bp	
NOV26a, CG147351-01 DNA Sequence	CACAAGAACAATATGCAGCTGAGATGAGTAAAGCTATTGCTTTTGAGATCATTCAGAAATACGAGCC TATCGAAGAAGTTAGGAAAGCACACCAAATGTCATTAGAAGGTTTTACAAGATACATGGATTCACGT GAATGTCTACTGTTTAAAAATGAATGTAGAAAAGTTTATCAAGATATGACTCATCCATTAAATGATT ATTTTATTTCATCTTCACATAACACATATTTGGTATCTGATCAATTATTGGGACCAAGTGACCTTTG GGGATATGTAAGTGCCCTTGTGAAAGGATGCCGTTGTTTGGAGATTGACTGCTGGGATGGAGCACAA AATGAACCTGTTGTATATCATGGCTACACACTCACAAGCAAACTTCTGTTTAAAACTGTTATCCAAG CTATACACAAGTATGCATTCATGGTGGCTTTAAATTTCCAGACCCCTGGTCTGCCCATGGATCTGCA AAATGGGAAATTTTTGGATAATGGTGGTTCTGGATATATTTTGAAACCACATTTCTTAAGAGAGAGT AAATCATACTTTAACCCAAGTAACATAAAAGAGGGTATGCCAATTACACTTACAATAAGGCTCATCA GTGGTATCCAGTTGCCTCTTACTCATTCATCATCTAACAAAGGTGATTCATTAGTAATTATAGAAGT TTTTGGTGTTCCAAATGATCAAATGAAGCAGCAGACTCGTGTAATTAAAAAAAATGCTTTTAGTCCA AGATGGAATGAAACATTCACATTTATTATTCATGTCCCAGAATTGGCATTGATACGTTTTGTTGTTG AAGGTCAAGGTTTAATAGCAGGAAATGAATTTCTTGGGCAATATACTTTGCCACTTCTATGCATGAA CAAAGGTTATCGTCGTATTCCTCTGTTTTCCAGAATGGGTGAGAGCCTTGAGCCTGCTTCACTGTTT GTTTATGTTTGGTACGTCAGATAACAGCTAAG

	ORF Start: ATG at 24		ORF Stop: TAA at 960

	SEQ ID NO: 120	312 aa	MW at 35720.0kD
NOV26a, CG147351-01 Protein Sequence	MSKAIAFEIIQKYEPIEEVRKAHQMSLEGFTRYMDSRECLLFKNECRKVYQDMTHPLNDYFISSSHN TYLVSDQLLGPSDLWGYVSALVKGCRCLEIDCWDGAQNEPVVYHGYTLTSKLLFKTVIQAIHKYAFM VALNFQTPGLPMDLQNGKFLDNGGSGYILKPHFLRESKSYFNPSNIKEGMPITLTIRLISGIQLPLT HSSSNKGDSLVIIEVFGVPNDQMKQQTRVIKKNAFSPRWNETFTFIIHVPELALIRFVVEGQGLIAG NEFLGQYTLPLLCMNKGYRRIPLFSRMGESLEPASLFVYVWYVR		

Further analysis of the NOV26a protein yielded the following properties shown in Table 26B.

Table 26B. Protein Sequence Properties NOV26a

PSort analysis:	0.5844 probability located in microbody (peroxisome); 0.1814 probability located in lysosome (lumen); 0.1000 probability located in mitochondrial matrix space; 0.0000 probability located in endoplasmic reticulum (membrane)
SignalP analysis:	No Known Signal Sequence Predicted

A search of the NOV26a protein against the Geneseq database, a proprietary database that contains sequences published in patents and patent publication, yielded several homologous proteins shown in Table 26C.

Table 26C. Geneseq Results for NOV26a

Geneseq Identifier	Protein/Organism/Length [Patent #, Date]	NOV26a Residues/ Match Residues	Identities/ Similarities for the Matched Region	Expect Value
AAU76817	Human phospholipase C 16839 polypeptide - Homo sapiens, 608 aa. [WO200206302-A2, 24-JAN-2002]	134..312 430..608	179/179 (100%) 179/179 (100%)	e-101
ABB90425	Human polypeptide SEQ ID NO 2801 - Homo sapiens, 179 aa. [WO200190304-A2, 29-NOV-2001]	134..312 1..179	179/179 (100%) 179/179 (100%)	e-101

AAU87271	Novel central nervous system protein #181 – Homo sapiens, 254 aa. [WO200155318-A2, 02-AUG-2001]	134..312 76..254	179/179 (100%) 179/179 (100%)	e-101
AAM95867	Human reproductive system related antigen SEQ ID NO: 4525 – Homo sapiens, 254 aa. [WO200155320-A2, 02-AUG-2001]	134..312 76..254	178/179 (99%) 178/179 (99%)	e-100
AAU22938	Novel human enzyme polypeptide #24 – Homo sapiens, 254 aa. [WO200155301-A2, 02-AUG-2001]	134..312 76..254	178/179 (99%) 178/179 (99%)	e-100

In a BLAST search of public sequence datbases, the NOV26a protein was found to have homology to the proteins shown in the BLASTP data in Table 26D.

5

Table 26D. Public BLASTP Results for NOV26a				
Protein Accession Number	Protein/Organism/Length	NOV26a Residues/ Match Residues	Identities/ Similarities for the Matched Portion	Expect Value
BAC05152	CDNA FLJ40406 fis, clone TESTI2037534, weakly similar to 1-PHOSPHATIDYLINOSITOL-4,5-B ISPHOSPHATE PHOSPHODIESTERASE DELTA 1 (EC 3.1.4.11) – Homo sapiens (Human), 390 aa.	134..312 212..390	179/179 (100%) 179/179 (100%)	e-101
Q96J70	Testis-development related NYD-SP27 – Homo sapiens (Human), 504 aa.	134..312 326..504	178/179 (99%) 178/179 (99%)	e-100
Q95JS0	Hypothetical 74.4 kDa protein – Macaca fascicularis (Crab eating macaque) (Cynomolgus monkey), 640 aa.	134..312 462..640	172/179 (96%) 177/179 (98%)	2e-97
Q95JS1	Hypothetical 74.6 kDa protein – Macaca fascicularis (Crab eating macaque) (Cynomolgus monkey), 641 aa.	134..312 463..641	172/179 (96%) 177/179 (98%)	2e-97
AAM95914	PLC-zeta – Mus musculus (Mouse), 647 aa.	134..312 467..646	135/181 (74%) 158/181 (86%)	7e-73

PFam analysis predicts that the NOV26a protein contains the domains shown in the Table 26E.

5

Table 26E. Domain Analysis of NOV26a			
Pfam Domain	NOV26a Match Region	Identities/ Similarities for the Matched Region	Expect Value
PI-PLC-X	52..133	45/83 (54%) 66/83 (80%)	4.3e-36
PI-PLC-Y	134..169	25/43 (58%) 33/43 (77%)	2.9e-17
C2	188..276	33/97 (34%) 73/97 (75%)	4.9e-20

Example 27.

The NOV27 clone was analyzed, and the nucleotide and encoded polypeptide
10   sequences are shown in Table 27A.

Table 27A. NOV27 Sequence Analysis			
	SEQ ID NO: 121	3136 bp	
NOV27a, CG147419-01 DNA Sequence	AGGGAGTCGTGTCGGCGCCACCCCGGCCCCCGAGCCCGCAGATTGCCCACCGAAGCTCGTGTGTGCA CCCCCGATCCCGCCAGCCACTCGCCCCTGGCCTCGCGGGCCGTGTCTCCGGCATCATGTGTGGTATA TTTGCTTACTTAAACTACCATGTTCCTCGAACGAGACGAGAAATCCTGGAGACCCTAATCAAAGGCC TTCAGAGACTGGAGTACAGAGGATATGATTCTGCTGGTGTGGGATTTGATGGAGGCAATGATAAAGA TTGGGAAGCCAATGCCTGCAAAACCCAGCTTATTAAGAAGAAAGGAAAAGTTAAGGCACTGGATGAA GAAGTTCACAAGCAACAAGATATGGATTTGGATATAGAATTTGATGTACACCTTGGAATAGCTCATA CCCGTTGGGCAACACATGGAGAACCCAGTCCTGTCAATAGCCACCCCCAGCGCTCTGATAAAAATAA TGAATTTATCGTTATTCACAATGGCATCATCACCAACTACAAAGACTTGAAAAAGTTTTTGGAAAGC AAAGGCTATGACTTCGAATCTGAAACAGACACAGAGACAATTGCCAAGCTCGTTAAGTATATGTATG ACAATCGGGAAAGTCAAGATACCAGCTTTACTACCTTGGTGGAGAGAGTTATCCAACAATTGGAAGG TGCTTTTGCACTTGTGTTTAAAAGTGTTCATTTTCCCGGGCAAGCAGTTGGCACAAGGCGAGGTAGC CCTCTGTTGATTGGTGTACGGAGTGAACATAAACTTTCTACTGATCACATTCCTATACTCTACAGAA CAGCTAGGACTCAGATTGGATCAAAATTCACACGGTGGGGATCACAGGGAGAAAGAGGCAAAGACAA GAAAGGAAGCTGCAATCTCTCTCGTGTGGACAGCACAACCTGCCTTTTCCCGGTGGAAGAAAAAGCA GTGGAGTATTACTTTGCTTCTGATGCAAGTGCTGTCATAGAACACACCAATCGCGTCATCTTTCTGG AAGATGATGATGTTGCAGCAGTAGTGGATGGACGTCTTTCTATCCATCGAATTAAACGAACTGCAGG AGATCACCCCGGACGAGCTGTGCAAACACTCCAGATGGAACTCCAGCAGATCATGAAGGGCAACTTC AGTTCATTTATGCAGAAGGAAATATTTGAGCAGCCAGAGTCTGTCGTGAACACAATGAGAGGAAGAG TCAACTTTGATGACTATACTGTGAATTTGGGTGGTTTGAAGGATCACATAAAGGAGATCCAGAGATG CCGGCGTTTGATTCTTATTGCTTGTGGAACAAGTTACCATGCTGGTGTAGCAACACGTCAAGTTCTT GAGGAGCTGACTGAGTTGCCTGTGATGGTGGAACTAGCAAGTGACTTCCTGGACAGAAACACACCAG TCTTTCGAGATGATGTTTGCTTTTTTCCTTAGTCAATCAGGTGAGACAGCAGATACTTTGATGGGTCT TCGTTACTGTAAGGAGAGAGGAGCTTTAACTGTGGGGATCACAAACACAGTTGGCAGTTCCATATCA CGGGAGACAGATTGTGGAGTTCATATTAATGCTGGTCCTGAGATTGGTGTGGCCAGTACAAAGGCTT ATACCAGCCAGTTTGTATCCCTTGTGATGTTTGCCCTTATGATGTGTGATGATCGGATCTCCATGCA AGAAAGACGCAAAGAGATCATGCTTGGATTGAAACGGCTGCCTGATTTGATTAAGGAAGTACTGAGC ATGGATGACGAAATTCAGAAACTAGCAACAGAACTTTATCATCAGAAGTCAGTTCTGATAATGGGAC GAGGCTATCATTATGCTACTTGTCTTGAAGGGGCACTGAAAATCAAAGAAATTACTTATATGCACTC TGAAGGCATCCTTGCTGGTGAATTGAAACATGGCCCTCTGGCTTTGGTGGATAAATTGATGCCTGTG

	ATCATGATCATCATGAGAGATCACACTTATGCCAAGTGTCAGAATGCTCTTCAGCAAGTGGTTGCTC GGCAGGGGCGGCCTGTGGTAATTTGTGATAAGGAGGATACTGAGACCATTAAGAACACAAAAGAAC GATCAAGGTGCCCCACTCAGTGGACTGCTTGCAGGGCATTCTCAGCGTGATCCCTTTACAGTTGCTG GCTTTCCACCTTGCTGTGCTGAGAGGCTATGATGTTGATTTCCCACGGAATCTTGCCAAATCTGTGA CTGTAGAGTGAGGAATATCTATACAAAATGTACGAAACTGTATGATTAAGCAACACAAGACACCTTT TGTATTTAAAACCTTGATTTAAAAATATCACCCCTTGAAGCCTTTTTTTAGTAAATCCTTATTTATAT ATCAGTTATAATTATTCCACTCAATATGTGATTTTTGTGAAGTTACCTCTTACATTTTCCCAGTAAT TTGTGGAGGACTTTGAATAATGGAATCTATATTGGAATCTGTATCAGAAAGATTCTAGCTATTATTT TCTTTAAAGAATGCTGGGTGTTGCATTTCTGGACCCTCCACTTCAATCTGAGAAGACAATATGTTTC TAAAAATTGGTACTTGTTTCACCATACTTCATTCAGACCAGTGAAAGAGTAGTGCATTTAATTGGAG TATCTAAAGCCAGTGGCAGTGTATGCTCATACTTGGACAGTTAGGGAAGGGTTTGCCAAGTTTTAAG AGAAGATGTGATTTATTTTGAAATTTGTTTCTGTTTTGTTTTTAAATCAAACTGTAAAACTTAAAAC TGAAAAATTTTATTGGTAGGATTTATATCTAAGTTTGGTTAGCCTTAGTTTCTCAGACTTGTTGTCT ATTATCTGTAGGTGGAAGAAATTTAGGAAGCGAAATATTACAGTAGTGCATTGGTGGGTCTCAATCC TTAACATATTTGCACAATTTTATAGCACAAACTTTAAATTCAAGCTGCTTTGGACAACTGACAATAT GATTTTAAATTTGAAGATGGGATGTGTGTACATGTTGGGTATCCTACTACTTTGTGTTTTCATCTCCTA AAAGTGTTTTTTTATTTCCTTGTATCTGTAGTCTTTTATTTTTTAAATGACTGCTGAATGACATATTT TATCTTGTTCTTTAAAATCACAACACAGAGCTGCTATTAAATTAATATTGATAT
	ORF Start: ATG at 123 ORF Stop: TGA at 2220

	SEQ ID NO: 122	699 aa	MW at 78793.6kD
NOV27a, CG147419-01 Protein Sequence	MCGIFAYLNYHVPRTRREILETLIKGLQRLEYRGYDSAGVGFDGGNDKDWEANACKTQLIKKKGKVK ALDEEVHKQQDMDLDIEFDVHLGIAHTRWATHGEPSPVNSHPQRSDKNNEFIVIHNGIITNYKDLKK FLESKGYDFESETDTETIAKLVKYMYDNRESQDTSFTTLVERVIQQLEGAFALVFKSVHFPGQAVGT RRGSPLLIGVRSEHKLSTDHIPILYRTARTQIGSKFTRWGSQGERGKDKKGSCNLSRVDSTTCLFPV EEKAVEYYFASDASAVIEHTNRVIFLEDDDVAAVVDGRLSIHRIKRTAGDHPGRAVQTLQMELQQIM KGNFSSFMQKEIFEQPESVVNTMRGRVNFDDYTVNLGGLKDHIKEIQRCRRLILIACGTSYHAGVAT RQVLEELTELPVMVELASDFLDRNTPVFRDDVCFFLSQSGETADTLMGLRYCKERGALTVGITNTVG SSISRETDCGVHINAGPEIGVASTKAYTSQFVSLVMFALMMCDDRISMQERRKEIMLGLKRLPDLIK EVLSMDDEIQKLATELYHQKSVLIMGRGYHYATCLEGALKIKEITYMHSEGILAGELKHGPLALVDK LMPVIMIIMRDHTYAKCQNALQQVVARQGRPVVICDKEDTETIKNTKRTIKVPHSVDCLQGILSVIP LQLLAFHLAVLRGYDVDFPRNLAKSVTVE

Further analysis of the NOV27a protein yielded the following properties shown in Table 27B.

10

Table 27B. Protein Sequence Properties NOV27a	
PSort analysis:	0.4902 probability located in mitochondrial inner membrane; 0.4400 probability located in plasma membrane; 0.3000 probability located in microbody (peroxisome); 0.2000 probability located in endoplasmic reticulum (membrane)
SignalP analysis:	No Known Signal Sequence Predicted

A search of the NOV27a protein against the Geneseq database, a proprietary

15 database that contains sequences published in patents and patent publication, yielded several homologous proteins shown in Table 27C.

Table 27C. Geneseq Results for NOV27a

Geneseq Identifier	Protein/Organism/Length [Patent #, Date]	NOV27a Residues/ Match Residues	Identities/ Similarities for the Matched Region	Expect Value
ABB05747	Human GFAT1L protein SEQ ID NO:1 - Homo sapiens, 699 aa. [WO200196574-A1, 20-DEC-2001]	1..699 1..699	698/699 (99%) 698/699 (99%)	0.0
AAY90260	Human GFAT protein sequence - Homo sapiens, 681 aa. [WO200037617-A1, 29-JUN-2000]	1..699 1..681	681/699 (97%) 681/699 (97%)	0.0
AAR43348	Human GFAT - Homo sapiens, 681 aa. [WO9321330-A, 28-OCT-1993]	1..699 1..681	680/699 (97%) 680/699 (97%)	0.0
AAY90261	Human GFAT II protein sequence - Homo sapiens, 682 aa. [WO200037617-A1, 29-JUN-2000]	1..699 1..682	541/701 (77%) 618/701 (87%)	0.0
AAW37772	Huma glutamine:fructose-6-phosph ate amidotransferase TGC028-4 - Homo sapiens, 682 aa. [EP824149-A2, 18-FEB-1998]	1..699 1..682	541/701 (77%) 618/701 (87%)	0.0

5   In a BLAST search of public sequence datbases, the NOV27a protein was found to

have homology to the proteins shown in the BLASTP data in Table 27D.

Table 27D. Public BLASTP Results for NOV27a

Protein Accession Number	Protein/Organism/Length	NOV27a Residues/ Match Residues	Identities/ Similarities for the Matched Portion	Expect Value
Q99MJ4	Glutamine: fructose-6-phosphate amidotransferase 1 muscle isoform GFAT1M - Mus musculus (Mouse), 697 aa.	1..699 1..697	688/699 (98%) 690/699 (98%)	0.0

A45055	glutamine--fructose-6-phosphate transaminase (isomerizing) (EC 2.6.1.16) - human, 681 aa.	1..699 1..681	681/699 (97%) 681/699 (97%)	0.0
Q06210	Glucosamine--fructose-6-phospha te aminotransferase [isomerizing] 1 (EC 2.6.1.16) (Hexosephosphate aminotransferase 1) (D-fructose-6- phosphate amidotransferase 1) (GFAT 1) (GFAT1) - Homo sapiens (Human), 680 aa.	2..699 1..680	680/698 (97%) 680/698 (97%)	0.0
BAB31882	Gfpt1 protein - Mus musculus (Mouse), 681 aa.	1..699 1..681	674/699 (96%) 676/699 (96%)	0.0
P47856	Glucosamine--fructose-6-phospha te aminotransferase [isomerizing] 1 (EC 2.6.1.16) (Hexosephosphate aminotransferase 1) (D-fructose-6- phosphate amidotransferase 1) (GFAT 1) (GFAT1) - Mus musculus (Mouse), 680 aa.	2..699 1..680	673/698 (96%) 675/698 (96%)	0.0

PFam analysis predicts that the NOV27a protein contains the domains shown in the Table 27E.

5

Table 27E. Domain Analysis of NOV27a			
Pfam Domain	NOV27a Match Region	Identities/ Similarities for the Matched Region	Expect Value
GATase_2	2..210	91/219 (42%) 202/219 (92%)	4.6e-127
SIS	378..512	52/156 (33%) 118/156 (76%)	2.2e-48
SIS	549..685	52/156 (33%) 124/156 (79%)	3.3e-46

Example 28.

10   The NOV28 clone was analyzed, and the nucleotide and encoded polypeptide sequences are shown in Table 28A.

Table 28A. NOV28 Sequence Analysis			
	SEQ ID NO: 123	2521 bp	
NOV28a, CG148102-01 DNA Sequence	ACTCTGCCCGACTCAGGGCTCCAGCGTGACATGGCTGAAGCGCACCAGGCCGTGGGCTTCCGACCCT CGCTGACCTCGGACGGGGCTGAAGTGGAACTCAGTGCCCCTGTGCTGCAGGAGATCTACCTCTCTGG CCTGCGCTCCTGGAAAAGGCATCTCTCACGTTTCTGGGTGCAGAATGACTTTCTCACCGGTGTGTTT CCTGCCAGCCCCCTCAGTTGGCTTTTCCTCTTCAGTGCCATCCAGCTTGCCTGGTTCCTCCAGCTGG ATCCTTCCTTAGGACTGATGGAGAAGATCAAAGAGTTGCTGCGGGGGGTCCTGGCAGCCGCGCTGTT TGCCTCGTGTTTGTGGGGAGCCCTGATCTTCACACTGCACGTGGCCCTGAGGCTGCTTCTGTCCTAC CACGGCTGGCTTCTTGAGCCCCACGGAGCCATGTCCTCCCCCACCAAGACCTGGCTGGCCCTGGTCC GCATCTTCTCTGGCCGCCACCCGATGCTGTTCAGTTACCAGCGCTCCCTGCCACGCCAGCCCGTGCC CTCTGTGCAGGACACCGTGCGCAAGTACCTGGAGTCGGTCCGGCCCATCCTCTCCGACGAGGACTTC GACTGGACCGCGGTCCTGGCGCAGGAATTCCTGAGGCTGCAGGCGTCGCTGCTGCAGTGGTACCTGC GGCTCAAGTCCTGGTGGGCGTCCAATTATGTGAGTGACTGGTGGGAGGAATTTGTGTACCTGCGCTC CCGAAATCCGCTGATGGTGAACAGCAATTACATGATGGACTTCCTGTATGTCACACCCACGCCT CTGCAGGCAGCTCGCGCTGGGAATGCCGTCCATGCCCTCCTCCTGTACCGCCACCGCCTGAACCGCC AGGAGATACCCCGGTGAGACTGATGGGAATGCGCCCCTTATGCTCTGCCCAGTACGAGAAGATCTT CAACACCACGCGGATTCCAGGGGTCCAAAAAGGTGAGACCATCCGCCACCTCCATGACAGCCAACAC GTGGCTGTCTTCCACCGGGGCCGATTCTTCCGCATGGGGACCCACTCCCGAAACAGCCTGCTTTCCC CGAGAGCCCTGGAGCAGCAGTTTCAGAGAATCCTGGATGATCCCTCACCGGCCTGCCCCCACGAGGA ACATCTGGCAGCTCTGACAGCTGCTCCCAGGGGCACGTGGGCCCAGGTGCGGACATCCCTGAAGACC CAGGCAGCGGAGGCCCTGGAGGCGGTGGAAGGGGCCGCTTTCTTTGTGTCACTGGATGCTGAGCCCG CGGGGCTCACCAGGGAGGACCCGGCAGCGTCGTTGGATGCCTACGCCCATGCTCTGCTGGCCGGCCG GGGCCATGATCGGTGGTTTGACAAATCCTTCACCCTAATCGTCTTCTCTAACGGGAAGCTGGGCCTC AGCGTGGAGCACTCCTGGGCCGACTGCCCCATCTCAGGACACATGTGGGAGTTCACTCTGGCTACAG AATGCTTTCAGCTGGGCTACTCAACAGACGGCCACTGCAAGGGGCACCCGGACCCCACACTACCCCA GCCCCAGCGGCTGCAATGGGACCTTCCAGACCAGGTGAGGCTGGGTATCTCTCTAGCCCTGAGGGGA GCCAAGATCTTGTCTGAAAATGTCGACTGCCATGTCGTTCCATTCTCCCCTATTTGGCAAGAGCTTCA TCCGACGCTGCCACCTCTCTTCAGACAGCTTCATCCAGATCGCCTTGCAACTGGCCCACTTCCGGGA CCCACAGTGCCTCGCCCTGTTCCGCGTGGCAGTGGACAAGCACCAGGCTCTGCTGAAGGCAGCCATG AGCGGGCAGGGAGTTGACCGCCACCTGTTTGCGCTGTACATCGTGTCCCGATTCCTCCACCTGCAGT CGCCCTTCCTGACCCAGGTCCATTCGGAGCAGTGGCAGCTGTCCACCAGCCAGATCCCTGTTCAGCA AATGCATCTGTTTGACGTCCACAATTACCCGGACTATGTTTCCTCAGGCGGTGGATTCGGGCCTGCT GATGACCATGGTTATGGTGTTTCTTATATCTTCATGGGGGATGGCATGATCACCTTCCACATCTCCA GCAAAAAATCAAGCACAAAAACGGATTCCCACAGGCTGGGGCAGCACATTGAGGACGCACTGCTGGA TGTGGCCTCCCTGTTCCAGGCGGACAGCATTTTAAGCGCCGGTTCAGAGGGTCAGGGAAGGAGAAC TCCAGGCACAGGTGTGGATTTCTCTCCCGCCAGACTGGGGCCTCCAAGGCCTCAATGACATCCACCG ACTTC**TGA**CTCCTTCCAGCAGGCAGCTGGCCTCTCCAAGGAATAAGGGTGAAATTGCCACAGCTGGC TGACACAGGACAGGGGCAACTGGTTTGGCAACCCCACATCCAGGCCAATAAAGATGTGTGAGCTGGG TGTGTGGTGTCTGCTATGCTCTTGGGCAGGGCAGGGGTAGAAGAGGTAAGGACCAGGGTGGAGGAGG ACAGAAGCTCCCATCCATTCCCAGGCCCAGCCAGGGATTCCC		
	ORF Start: ATG at 31		ORF Stop: TGA at 2284

	SEQ ID NO: 124	751 aa	MW at 84918.2kD
NOV28a, CG148102-01 Protein Sequence	MAEAHQAVGFRPSLTSDGAEVELSAPVLQEIYLSGLRSWKRHLSRFWVQNDFLTGVFPASPLSWLFL FSAIQLAWFLQLDPSLGLMEKIKELLRGVLAAALFASCLWGALIFTLHVALRLLLSYHGWLLEPHGA MSSPTKTWLALVRIFSGRHPMLFSYQRSLPRQPVPSVQDTVRKYLESVRPILSDEDFDWTAVLAQEF LRLQASLLQWYLRLKSWWASNYVSDWWEEFVYLRSRNPLMVNSNYYMMDFLYVTPTPLQAARAGNAV HALLLYRHRLNRQEIPPVRLMGMRPLCSAQYEKIFNTTRIPGVQKGETIRHLHDSQHVAVFHRGRFF RMGTHSRNSLLSPRALEQQFQRILDDPSPACPHEEHLAALTAAPRGTWAQVRTSLKTQAAEALEAVE GAAFFVSLDAEPAGLTREDPAASLDAYAHALLAGRGHDRWFDKSFTLIVFSNGKLGLSVEHSWADCP ISGHMWEFTLATECFQLGYSTDGHCKGHPDPTLPQPQRLQWDLPDQVRLGISLALRGAKILSENVDC HVVPFSLFGKSFIRRCHLSSDSFIQIALQLAHFRDPQCLALFRVAVDKHQALLKAAMSGQGVDRHLF ALYIVSRFLHLQSPFLTQVHSEQWQLSTSQIPVQQMHLFDVHNYPDYVSSGGGFGPADDHGYGVSYI FMGDGMITFHISSKKSSTKTDSHRLGQHIEDALLDVASLFQAGQHFKRRFRGSGKENSRHRCGFLSR QTGASKASMTSTDF		

	SEQ ID NO: 125	2748 bp	

NOV28b, CG148102-02 DNA Sequence	CGAGAGACAGGAATCGGGGTTTCTGGGTGACGGTGATCTCGGGGTGGGCAGGACTCCAAAGGCCCGT
	CGACCCGGTGGTGGACTCCTTGCACTGGGATTGGACATATGCAAGCGGGAGATTTGGGGCCGGCGCT
	CAAAATCGGGGGGCGGGGGTGGACTCGGGTTTGGACCCCAGGATCCGATCAGCGGACCCTTGATTCA
	ACGTGGGCTCCAGCGTGACATGGCTGAAGCGCACCAGGCCGTGGGCTTCCGACCCTCGCTGACCTCG
	GACGGGGCTGAAGTGGAACTCAGTGCCCCTGTGCTGCAGGAGATCTACCTCTCTGGCCTGCGCTCCT
	GGAAAAGGCATCTCTCACGTTTCTGGAATGACTTTCTCACCGGTGTGTTTCCTGCCAGCCCCCTCAG
	TTGGCTTTTCCTCTTCAGTGCCATCCAGCTTGCCTGGTTCCTCCAGCTGGATCCTTCCTTAGGACTG
	ATGGAGAAGATCAAAGAGTTGCTGCCTGACTGGGGTGGACAACACCACGGGCTCCGGGGGGTCCTGG
	CAGCCGCGCTGTTTGCCTCGTGTTTGTGGGGAGCCCTGATCTTCACACTGCACGTGGCCCTGAGGCT
	GCTTCTGTCCTACCACGGCTGGCTTCTTGAGCCCCACGGAGCCATGTCCTCCCCCACCAAGACCTGG
	CTGGCCCTGGTCCGCATCTTCTCTGGCCGCCACCCGATGCTGTTCAGTTACCAGCGCTCCCTGCCAC
	GCCAGCCCGTGCCCTCTGTGCAGGACACCGTGCGCAAGTACCTGGAGTCGGTCCGGCCCATCCTCTC
	CGACGAGGACTTCGACTGGACCGCGGTCCTGGCCGCAGGAATTCCTGAGGCTGCAGGCGTCACTGCTG
	CAGTGGTACCTGCGGCTCAAGTCCTGGTGGGCGTCCAATTATGTCAGTGACTGGTGGGAGGAATTTG
	TGTACCTGCGCTCCCGAAATCCGCTGATGGTGAACAGCAACTATTACATGATGGACTTCCTGTATGT
	CACACCCACGCCTCTGCAGGCAGCTCGCGCTGGGAATGCCGTCCATGCCCTCCTCCTGTACCGCCAC
	CGCCTGAACCGCCAGGAGATACCCCGACTTTGCTGATGGGAATGCGCCCCCTTATGCTCTGCCCAGT
	ACGAGAAGATCTTCAACACCACGCGGATTCCAGGGGTCCAAAAAGACTACATCCGCCACCTCCATGA
	CAGCCAACACGTGGCTGTCTTCCACCGGGGCCGATTCTTCCGCATGGGGACCCACTCCCGAAACAGC
	CTGCTTTCCCCGAGAGCCCTGGAGCAGCAGTTTCAGAGAATCCTGGATGATCCCTCACCGGCCTGCC
	CCCACGAGGAACATCTGGCAGGTCTGACAGCTGCTCCCAGGGCACGTGGGCCCAGGTGCGGACATC
	CCTGAAGACCCAGGCAGCGGAGGCCCTGGAGGCGGTGGAAGGGGCCGCTTTCTTTGTGTCACTGGAT
	GCTGAGCCCGCGGGGCTCACCAGGGAGGACCCGGCAGCGTCGTTGGATGCCTACGCCCATGCTCTGC
	TGGCTGGCCGGGGCCATGATCGCTGGTTTGACAAATCCTTCACCCTAATCGTCTTCTCTAACGGGAA
	GCTGGGCCTCAGCGTGGAGCACTCCTGGGCCGACTGCCCCATCTCAGGACACATGTGGGAGTTCACT
	CTGGCTACAGAATGCTTTCAGCTGGGCTACTCAACAGATGGCCACTGCAAGGGGCACCCGGACCCCA
	CACTACCCCAGCCCCAGCGGCTGCAATGGGACCTTCCAGACCAGATCCACTCCTCCATCTCTCTAGC
	CCTGAGGGGAGCCAAGATCTTGTCTGAAAATGTCGACTGCCATGTCGTTCCATTCTCCCTATTTGGC
	AAGAGCTTCATCCGACGCTGCCACCTCTCTTCAGACAGCTTCATCCAGATCGCCTTGCAACTGGCCC
	ACTTCCGGGACAGGGGTCAATTCTGCCTGACTTATGAGTCGGCCATGACTCGCTTATTCCTGGAAGG
	CCGGACGGAGACGGTGCGGTCTTGCACGAGGGAGGCCTGCAACTTTGTCAGGGCCATGGAGGACAAA
	GAGAAGACGGACCCACAGTGCCTCGCCCTGTTCCGCGTGGCAGTGGACAAGCACCAGGCTCTGCTGA
	AGGCAGCCATGAGCGGGCAGGGAGTTGACCGCCACCTGTTTGCGCTGTACATCGTGTCCCGATTCCT
	CCACCTGCAGTCGCCCTTCCTGACCCAGGTCCATTCGGAGCAGTGGCAGCTGTCCACCAGCCAGATC
	CCTGTTCAGCAAATGCATCTGTTTGACGTCCACAATTACCCGGACTATGTTTCCTCAGGCGGTGGAT
	TCGGGCCTGCTGATGACCATGGTTATGGTGTTTCTTATATCTTCATGGGGGATGGCATGATCACCTT
	CCACATCTCCAGCAAAAAATCAAGCACAAAAACGGATTCCCACAGGCTGGGGCAGCACATTGAGGAC
	GCACTGCTGGATGTGGCCTCCCTGTTCCAGGCGGGACAGCATTTTAAGCGCCGGTTCAGAGGGTCAG
	GGAAGGAGAACTCCAGGCACAGGTGTGGATTTCTCTCCCGCCAGACTGGGGCGTCCAAGGCCTCAAT
	GACATCCACCGACTTCTGACTCCTTCCAGCAGGCAGCTGGCCTCTCCAAGGAATAAGGGTGAAATTG
	CCACAGCTGGCTGACACAGGACAGGGGCAACTGGTTTGGCAACCCCACATCCAGGCAAATAAAGATG
	G
	ORF Start: ATG at 221		ORF Stop: TGA at 2630

	SEQ ID NO: 126	803 aa	MW at 90987.8kD
NOV28b, CG148102-02 Protein Sequence	MAEAHQAVGFRPSLTSDGAEVELSAPVLQEIYLSGLRSWKRHLSRFWNDFLTGVFPASPLSWLFLFS AIQLAWFLQLDPSLGLMEKIKELLPDWGGQHHGLRGVLAAALFASCLWGALIFTLHVALRLLLSYHG WLLEPHGAMSSPTKTWLALVRIFSGRHPMLFSYQRSLPRQPVPSVQDTVRKYLESVRPILSDEDFDW TAVLAQEFLRLQASLLQWYLRLKSWWASNYVSDWWEEFVYLRSRNPLMVNSNYYMMDFLYVTPTPLQ AARAGNAVHALLLYRHRLNRQEIPPTLLMGMRPLCSAQYEKIFNTTRIPGVQKDYIRHLHDSQHVAV FHRGRFFRMGTHSRNSLLSPRALEQQFQRILDDPSPACPHEEHLAALTAAPRGTWAQVRTSLKTQAA EALEAVEGAAFFVSLDAEPAGLTREDPAASLDAYAHALLAGRGHDRWFDKSFTLIVFSNGKLGLSVE HSWADCPISGHMWEFTLATECFQLGYSTDGHCKGHPDPTLPQPQRLQWDLPDQIHSSISLALRGAKI LSENVDCHVVPFSLFGKSFIRRCHLSSDSFIQIALQLAHFRDRGQFCLTYESAMTRLFLEGRTETVR SCTREACNFVRAMEDKEKTDPQCLALFRVAVDKHQALLKAAMSGQGVDRHLFALYIVSRFLHLQSPF LTQVHSEQWQLSTSQIPVQQMHLFDVHNYPDYVSSGGGFGPADDHGYGVSYIFMGDGMITFHISSKK SSTKTDSHRLGQHIEDALLDVASLFQAGQHFKRRFRGSGKENSRHRCGFLSRQTGASKASMTSTDF

Sequence comparison of the above protein sequences yields the following sequence relationships shown in Table 28B.

Table 28B. Comparison of NOV28a against NOV28b.		
Protein Sequence	**NOV28a Residues/ Match Residues**	**Identities/ Similarities for the Matched Region**
NOV28b	1..751 1..803	717/806 (88%) 719/806 (88%)

5   Further analysis of the NOV28a protein yielded the following properties shown in
Table 28C.

Table 28C. Protein Sequence Properties NOV28a	
PSort analysis:	0.7900 probability located in plasma membrane; 0.6400 probability located in microbody (peroxisome); 0.3000 probability located in Golgi body; 0.2000 probability located in endoplasmic reticulum (membrane)
SignalP analysis:	Cleavage site between residues 5 and 6

10

A search of the NOV28a protein against the Geneseq database, a proprietary
database that contains sequences published in patents and patent publication, yielded
several homologous proteins shown in Table 28D.

15

Table 28D. Geneseq Results for NOV28a				
Geneseq Identifier	**Protein/Organism/Length [Patent #, Date]**	**NOV28a Residues/ Match Residues**	**Identities/ Similarities for the Matched Region**	**Expect Value**
AAY79220	Human transferase TRNSFS-12 - Homo sapiens, 803 aa. [WO200014251-A2, 16-MAR-2000]	1..751 1..803	740/806 (91%) 742/806 (91%)	0.0
AAE10322	Human carnitine acyltransferase, 26886 - Homo sapiens, 803 aa. [WO200166759-A2, 13-SEP-2001]	1..751 1..803	739/806 (91%) 742/806 (91%)	0.0

AAW14438	Type I carnitine palmitoyl transferase-like protein - Homo sapiens, 772 aa. [JP09009969-A, 14-JAN-1997]	1..711 1..766	375/770 (48%) 495/770 (63%)	0.0
ABG04960	Novel human diagnostic protein #4951 - Homo sapiens, 521 aa. [WO200175067-A2, 11-OCT-2001]	224..571 92..471	337/381 (88%) 339/381 (88%)	0.0
ABB67527	Drosophila melanogaster polypeptide SEQ ID NO 29373 - Drosophila melanogaster, 780 aa. [WO200171042-A2, 27-SEP-2001]	1..717 1..765	315/775 (40%) 447/775 (57%)	e-161

In a BLAST search of public sequence datbases, the NOV28a protein was found to have homology to the proteins shown in the BLASTP data in Table 28E.

5

Table 28E. Public BLASTP Results for NOV28a				
Protein Accession Number	Protein/Organism/Length	NOV28a Residues/ Match Residues	Identities/ Similarities for the Matched Portion	Expect Value
Q8TCG5	Carnitine palmitoyltransferase IC - Homo sapiens (Human), 803 aa.	1..751 1..803	740/806 (91%) 742/806 (91%)	0.0
CAC88591	Sequence 1 from Patent WO0166759 - Homo sapiens (Human), 803 aa.	1..751 1..803	739/806 (91%) 742/806 (91%)	0.0
AAH29104	Similar to carnitine palmitoyltransferase IC - Homo sapiens (Human), 792 aa.	1..751 1..792	729/806 (90%) 731/806 (90%)	0.0
P32198	Carnitine O-palmitoyltransferase I, mitochondrial liver isoform (EC 2.3.1.21) (CPT I) (CPTI-L) - Rattus norvegicus (Rat), 773 aa.	1..710 1..765	394/768 (51%) 524/768 (67%)	0.0

| Q9BWK0 | Similar to carnitine palmitoyltransferase I, liver - Homo sapiens (Human), 756 aa. | 1..690
1..745 | 381/748 (50%)
510/748 (67%) | 0.0 |

PFam analysis predicts that the NOV28a protein contains the domains shown in the Table 28F.

5

Table 28F. Domain Analysis of NOV28a			
Pfam Domain	NOV28a Match Region	Identities/ Similarities for the Matched Region	Expect Value
Carn_acyltransf	162..708	208/680 (31%) 437/680 (64%)	1.5e-167

Example 29.

10   The NOV29 clone was analyzed, and the nucleotide and encoded polypeptide sequences are shown in Table 29A.

Table 29A. NOV29 Sequence Analysis			
	SEQ ID NO: 127	1776 bp	
NOV29a, CG148431-01 DNA Sequence	ACTAAAGCCTGCAGAGACCTCTGAAGGAAAACCTGTCCCGGGCTCTGTCACTTCACACCCATGGCTA ACCCTGGAGGTGGTGCTGTTTGCAACGGGAAACTTCACAATCACAAGAAACAGAGCAATGGCTCACA AAGCAGAAACTGCACAAAGAATGGAATAGTGAAGGAAGCCCAGCAAAATGGGAAGCCACATTTTTAT GATAAGCTCATTGTTGAATCGTTTGAGGAAGCACCCCTTCATGTTATGGTTTTCACTTACATGGGAT ATGGAATTGGAACCCTGTTTGGCTATCTCAGAGACTTTTTAAGAAACTGGGGAATAGAAAAATGCAA CGCAGCTGTGGAAAGAAAAGAACAAAAAGATTTTGTGCCACTGTATCAAGACTTTGAAAATTTTTAT ACAAGAAACCTTTACATGCGAATCAGAGACAACTGGAACCGGCCCATCTGCAGTGCCCCAGGGCCTC TGTTTGATTTGATGGAGAGGGTATCAGACGACTATAACTGGACGTTTAGGTTTACTGGAAGAGTCAT CAAAGATGTCATCAACATGGGCTCCTATAACTTCCTTGGTCTTGCAGCCAAGTATGATGAGTCTATG AGGACAATAAAGGATGTTTTAGAGGTGTATGGCACAGGCGTGGCCAGCACCAGGCATGAAATGGGCA CCTTGGATAAGCACAAGGAGTTGGAGGACCTTGTGGCTAAGTTCCTGAATGTGGAAGCAGCTATGGT CTTTGGGATGGGATTCGCAACTAACTCAATGAATATCCCAGCATTAGTTGGAAAGGGATGCCTCATT TTAAGTGATGAGTTAAACCACACATCGCTTGTGCTTGGGGCCCGACTCTCAGGTGCAACCATAAGAA TCTTCAAACACAACAACACACAAAGCCTAGAGAAGCTCCTGAGAGATGCTGTCATCTATGGCCAGCC TCGAACCCGCAGAGCTTGGAAAAAGATTCTCATCCTGGTGGAGGGTGTCTACAGCATGGAAGGTTCC ATCGTGCATCTGCCCCAGATCATAGCTCTAAAGAAGAAATACAAGGCTTACCTCTACATAGATGAAG CTCACAGTATTGGGGCCGTGGGCCCAACCGGCCGGGGTGTCACGGAGTTCTTTGGACTAGACCCTCA TGAAGTTGATGTGCTCATGGGCACATTCACCAAAAGTTTTGGAGCTTCAGGAGGTTACATAGCTGGA AGGAAGGACCTCGTGGATTATTTACGGGTTCACTCGCATAGTGCTGTTTATGCTTCATCCATGAGCC CACCGATAGCAGAGCAAATCATCAGATCACTAAAACTTATCATGGACTGGATGGGGACCACTCAAGG GCTGCAGAGAGTACAGCAACTTGCGAAAAACACAAGATACTTCAGACAAAGACTGCAGGAAATGGGA TTCATTATCTATGGCAATGAGAATGCTTCTGTTGTTCCTCTGCTTCTTTATATGCCTGGTAAAGTAG CGGCTTTTGCAAGGCATATGCTAGAGAAAAAAATTGGAGTGGTGGTCGTGGGATTTCCAGCCACTCC CCTCGCAGAAGCTCGGGCTCGGTTTTGTGTTTCAGCGGCACATACCCGGGAGATGTTAGACACGGTT TTAGAAGCTCTTGATGAAATGGGTGATCTCTTGCAACTGAAATATTCCCGGCACAAGAAGTCAGCAC GTCCTGAGCTCTATGATGAGACGAGCTTTGAACTCGAAGATTAAGTTTCCTGGTCCTGAATGACACA TAAAGACTTTGCGAGAAAGACCTCCCTCCTTGCC
	ORF Start: ATG at 61		ORF Stop: TAA at 1717

	SEQ ID NO: 128	552 aa	MW at 62048.9kD
NOV29a, CG148431-01 Protein Sequence	MANPGGGAVCNGKLHNHKKQSNGSQSRNCTKNGIVKEAQQNGKPHFYDKLIVESFEEAPLHVMVFTY MGYGIGTLFGYLRDFLRNWGIEKCNAAVERKEQKDFVPLYQDFENFYTRNLYMRIRDNWNRPICSAP GPLFDLMERVSDDYNWTFRFTGRVIKDVINMGSYNFLGLAAKYDESMRTIKDVLEVYGTGVASTRHE MGTLDKHKELEDLVAKFLNVEAAMVFGMGFATNSMNIPALVGKGCLILSDELNHTSLVLGARLSGAT IRIFKHNNTQSLEKLLRDAVIYGQPRTRRAWKKILILVEGVYSMEGSIVHLPQIIALKKKYKAYLYI DEAHSIGAVGPTGRGVTEFFGLDPHEVDVLMGTFTKSFGASGGYIAGRKDLVDYLRVHSHSAVYASS MSPPIAEQIIRSLKLIMGLDGTTQGLQRVQQLAKNTRYFRQRLQEMGFIIYGNENASVVPLLLYMPG KVAAFARHMLEKKIGVVVVGFPATPLAEARARFCVSAAHTREMLDTVLEALDEMGDLLQLKYSRHKK SARPELYDETSFELED		

	SEQ ID NO: 129	1492 bp	
NOV29b, CG148431-02 DNA Sequence	CACCGGATCCACCATGGCTAACCCTGGAGGTGGTGCTGTTTGCAACGGGAAACTTCACAATCACAAG AAACAGAGCAATGGCTCACAAAGCAGAAACTGCACAAAGAATGGAATAGTGAAGGAAGCCCAGGATT TTGTGCCACTGTATCAAGACTTTGAAAATTTTTATACAAGAAACCTTTACATGCGAATCAGAGACAA CTGGAACCGGCCCATCTGCAGTGCCCCAGGGCCTCTGTTTGATGTGATGGAGAGGGTATCGGACGAC TATAACTGGACGTTTAGGTTTACTGGAAGAGTCATCAAAGATGTCATCAACATGGGCTCCTATAACT TCCTTGGTCTTGCAGCCAAGTATGATGAGTCTATGAGGACAATAAAGGATGTTTTAGAGGTGTATGG CACAGGCGTGGCCAGCACCAGGCATGAAATGGGCACCTTGGATAAGCACAAGGAGTTGGAGGACCTT GTGGCTAAGTTCCTGAATGTGGAAGCAGCTATGGTCTTTGGGATGGGATTCGCAACTAACTCAATGA ATATCCCAGCATTAGTTGGAAAGGGATGCCTCATTTTAAGTGATGAGTTAAACCACACATCGCTTGT GCTTGGGGCCCGACTCTCAGGTGCAACCATAAGAATCTTCAAACACAACAACACACAAAGCCTAGAG AAGCTCCTGAGAGATGCTGTCATCTATGGCCAGCCTCGAACCCGCAGAGCTTGGAAAAAGATTCTCA TCCTGGTGGAGGGTGTCTACAGCATGGAAGGTTCCATCGTGCATCTGCCCCAGATCATAGCTCTAAA GAAGAAATACAAGGCTTACCTCTACATAGATGAAGCTCACAGTATTGGGGCCGTGGGCCCAACCGGC CGGGGTGTCACGGAGTTCTTTGGACTAGACCCTCATGAAGTTGATGTGCTCATGGGCACATTCACCA AAAGTTTTGGAGCTTCAGGAGGTTACATAGCTGGAAGGAAGGACCTCGTGGATTATTTACGGGTTCA CTCGCATAGTGCTGTTTATGCTTCATCCATGAGCCCACCGATAGCAGAGCAAATCATCAGATCACTA AAACTTATCATGGGACTGGATGGGACCACTCAAGGGCTGCAGAGAGTACAGCAACTTGCGAAAAACA CAAGATACTTCAGACAAAGACTGCAGGAAATGGGATTCATTATCTATGGCAATGAGAATGCTTCTGT TGTTCCTCTGCTTCTTTATATGCCTGGTAAAGTAGCGGCTTTTGCAAGGCATATGCTAGAGAAAAAA ATTGGAGTGGTGGTCGTGGGATTTCCAGCCACTCCCCTCGCAGAAGCTCGGGCTCGGTTTTGTGTTT CAGCGGCACATACCCGGGAGATGTTAGACACGGTTTTAGAAGCTCTTGATGAAATGGGTGATCTCTT GCAACTGAAATATTCCCGGCACAAGAAGTCAGCACGTCCTGAGCTCTATGATGAGACGAGCTTTGAA CTCGAAGATCTCGAGGGC		
	ORF Start: ATG at 14		ORF Stop: at 1484

	SEQ ID NO: 130	490 aa	MW at 54766.5kD
NOV29b, CG148431-02 Protein Sequence	MANPGGGAVCNGKLHNHKKQSNGSQSRNCTKNGIVKEAQDFVPLYQDFENFYTRNLYMRIRDNWNRP ICSAPGPLFDVMERVSDDYNWTFRFTGRVIKDVINMGSYNFLGLAAKYDESMRTIKDVLEVYGTGVA STRHEMGTLDKHKELEDLVAKFLNVEAAMVFGMGFATNSMNIPALVGKGCLILSDELNHTSLVLGAR LSGATIRIFKHNNTQSLEKLLRDAVIYGQPRTRRAWKKILILVEGVYSMEGSIVHLPQIIALKKKYK AYLYIDEAHSIGAVGPTGRGVTEFFGLDPHEVDVLMGTFTKSFGASGGYIAGRKDLVDYLRVHSHSA VYASSMSPPIAEQIIRSLKLIMGLDGTTQGLQRVQQLAKNTRYFRQRLQEMGFIIYGNENASVVPLL LYMPGKVAAFARHMLEKKIGVVVVGFPATPLAEARARFCVSAAHTREMLDTVLEALDEMGDLLQLKY SRHKKSARPELYDETSFELED		

Sequence comparison of the above protein sequences yields the following sequence relationships shown in Table 29B.

Table 29B. Comparison of NOV29a against NOV29b.		
Protein Sequence	NOV29a Residues/ Match Residues	Identities/ Similarities for the Matched Region
NOV29b	98..552 36..490	438/455 (96%) 440/455 (96%)

5   Further analysis of the NOV29a protein yielded the following properties shown in Table 29C.

Table 29C. Protein Sequence Properties NOV29a	
PSort analysis:	0.4761 probability located in microbody (peroxisome); 0.3000 probability located in nucleus; 0.2077 probability located in lysosome (lumen); 0.1000 probability located in mitochondrial matrix space
SignalP analysis:	No Known Signal Sequence Predicted

10

A search of the NOV29a protein against the Geneseq database, a proprietary database that contains sequences published in patents and patent publication, yielded several homologous proteins shown in Table 29D.

15

Table 29D. Geneseq Results for NOV29a				
Geneseq Identifier	Protein/Organism/Length [Patent #, Date]	NOV29a Residues/ Match Residues	Identities/ Similarities for the Matched Region	Expect Value
AAE22153	Human TRNFR-15 protein - Homo sapiens, 552 aa. [WO200226950-A2, 04-APR-2002]	1..552 1..552	551/552 (99%) 552/552 (99%)	0.0
AAG73598	Human colon cancer antigen protein SEQ ID NO:4362 - Homo sapiens, 391 aa. [WO200122920-A2, 05-APR-2001]	201..549 38..387	269/352 (76%) 316/352 (89%)	e-158
ABB60160	Drosophila melanogaster polypeptide SEQ ID NO 7272 - Drosophila melanogaster. 597 aa.	54..543 114..597	256/491 (52%) 350/491 (71%)	e-151

	[WO200171042-A2, 27-SEP-2001]			
AAE21820	Human serine palmitoyltransferase (SPT)-like enzyme #2 - Homo sapiens, 230 aa. [WO200224884-A2, 28-MAR-2002]	47..276 1..230	228/230 (99%) 230/230 (99%)	e-133
AAY32003	Rice serine palmitoyltransferase Lcb2 subunit - Oryza sativa, 489 aa. [WO9949021-A1, 30-SEP-1999]	59..541 5..483	237/485 (48%) 333/485 (67%)	e-133

In a BLAST search of public sequence datbases, the NOV29a protein was found to have homology to the proteins shown in the BLASTP data in Table 29E.

5

Table 29E. Public BLASTP Results for NOV29a				
Protein Accession Number	Protein/Organism/Length	NOV29a Residues/ Match Residues	Identities/ Similarities for the Matched Portion	Expect Value
Q9UGB6	DJ718P11.1.1 (Novel class II aminotransferase similar to serine palmotyltransferase (Isoform 1)) - Homo sapiens (Human), 414 aa (fragment).	102..515 1..414	414/414 (100%) 414/414 (100%)	0.0
O15270	Serine palmitoyltransferase 2 (EC 2.3.1.50) (Long chain base biosynthesis protein 2) (LCB 2) (Serine-palmitoyl-CoA transferase 2) (SPT 2) - Homo sapiens (Human), 562 aa.	7..549 18..558	383/546 (70%) 449/546 (82%)	0.0
P97363	Serine palmitoyltransferase 2 (EC 2.3.1.50) (Long chain base biosynthesis protein 2) (LCB 2) (Serine-palmitoyl-CoA transferase 2) (SPT 2) - Mus musculus (Mouse), 560 aa.	7..549 18..556	379/546 (69%) 449/546 (81%)	0.0
JC5180	serine C-palmitoyltransferase (EC 2.3.1.50) Lcb2 chain - mouse, 560 aa.	7..549 18..556	378/546 (69%) 449/546 (82%)	0.0

| O54694 | Serine palmitoyltransferase 2 (EC 2.3.1.50) (Long chain base biosynthesis protein 2) (LCB 2) (Serine-palmitoyl-CoA transferase 2) (SPT 2) - Cricetulus griseus (Chinese hamster), 560 aa. | 7..549 18..556 | 377/546 (69%) 446/546 (81%) | 0.0 |

PFam analysis predicts that the NOV29a protein contains the domains shown in the Table 29F.

Table 29F. Domain Analysis of NOV29a

Pfam Domain	NOV29a Match Region	Identities/ Similarities for the Matched Region	Expect Value
aminotran_1_2	193..521	71/363 (20%) 237/363 (65%)	2.6e-29

Example 30.

The NOV30 clone was analyzed, and the nucleotide and encoded polypeptide sequences are shown in Table 30A.

Table 30A. NOV30 Sequence Analysis

	SEQ ID NO: 131	576 bp	
NOV30a, CG148888-01 DNA Sequence	TGAGCCAGCCCCGGATGACCCTGCGACCTGGAACAATGCGGCTGGCCTGCATGTTCTCTTCCATCCT GCTGTTCGGAGCTGCAGGCCTCCTCCTCTTCATCAGCCTGCAGGACCCTACGGAGCTCGCCCCCCAG CAGGTGCCAGGAATAAAGTTCAACATCAGGCCAAGGCAGCCCCACCACGACCTCCCACCAGGCGGCT CTGGGGTGCGTTTTCCCGAGTTCGTCCAGTACCTGCTGGACGTGCACCGGCCCGTGGGGATGGACAT TCACTGGGACCATGTCAGCCGGCTCTGCAGCCCCTGCCTCATCGACTACGATTTCGTAGGCAAGTTC GAGAGCATGGAGGACGATGCCAACTTCTTCCTGAGCCTCATCCGCGCGCCGCGGAACCTGACCTTCC CCCGGTTCAAGGACCGGCACTCGCAGGAGGCGCGGGACCACAGCGAGGATCGCCCACCAGTACTTCGC CCAACTCTCGGCCCTGCAAAGGCAGCGCACCTACGACTTCTACTACATGGATTACCTGATGTTCAAC TATTCCAAGCCCTTTACAGATCTGTACTGAGGGGCGCCGC		
	ORF Start: ATG at 15		ORF Stop: TGA at 564

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	SEQ ID NO: 132	183 aa	MW at 21347.3kD
NOV30a, CG148888-01 Protein Sequence	MTLRPGTMRLACMFSSILLFGAAGLLLFISLQDPTELAPQQVPGIKFNIRPRQPHHDLPPGGSGVRF PEFVQYLLDVHRPVGMDIHWDHVSRLCSPCLIDYDFVGKFESMEDDANFFLSLIRAPRNLTFPRFKD RHSQEARTTARIAHQYFAQLSALQRQRTYDFYYMDYLMFNYSKPFTDLY		

5   Further analysis of the NOV30a protein yielded the following properties shown in Table 30B.

Table 30B. Protein Sequence Properties NOV30a	
PSort analysis:	0.8650 probability located in lysosome (lumen); 0.8200 probability located in outside; 0.3657 probability located in microbody (peroxisome); 0.1000 probability located in endoplasmic reticulum (membrane)
SignalP analysis:	Cleavage site between residues 38 and 39

10   A search of the NOV30a protein against the Geneseq database, a proprietary database that contains sequences published in patents and patent publication, yielded several homologous proteins shown in Table 30C.

15

Table 30C. Geneseq Results for NOV30a				
Geneseq Identifier	Protein/Organism/Length [Patent #, Date]	NOV30a Residues/ Match Residues	Identities/ Similarities for the Matched Region	Expect Value
ABB53266	Human polypeptide #6 - Homo sapiens, 424 aa. [WO200181363-A1, 01-NOV-2001]	62..183 303..424	121/122 (99%) 121/122 (99%)	4e-69
ABB53265	Human polypeptide #5 - Homo sapiens, 628 aa. [WO200181363-A1, 01-NOV-2001]	62..183 507..628	121/122 (99%) 121/122 (99%)	4e-69
AAE15437	Human drug metabolising enzyme (DME)-4 - Homo sapiens, 396 aa. [WO200179468-A2, 25-OCT-2001]	62..183 275..396	121/122 (99%) 121/122 (99%)	4e-69
AAB85083	Human interleukin-6 (IL-6) like polypeptide - Homo sapiens, 171 aa. [WO200142484-A1, 14-JUN-2001]	62..183 50..171	121/122 (99%) 121/122 (99%)	4e-69
AAM24429	Murine EST encoded protein SEQ ID NO: 1954 - Mus musculus, 424 aa. [WO200154477-A2, 02-AUG-2001]	62..183 303..424	121/122 (99%) 121/122 (99%)	4e-69

In a BLAST search of public sequence datbases, the NOV30a protein was found to have homology to the proteins shown in the BLASTP data in Table 30D.

Table 30D. Public BLASTP Results for NOV30a				
Protein Accession Number	Protein/Organism/Length	NOV30a Residues/ Match Residues	Identities/ Similarities for the Matched Portion	Expect Value
Q9H3N2	GalNAc 4-sulfotransferase (GalNAc-4-O-sulfotransferase 1) (Carbohydrate (N-acetylgalactosamine 4-0) sulfotransferase 8) (Hypothetical 48.8 kDa protein) - Homo sapiens (Human), 424 aa.	62..183 303..424	121/122 (99%) 121/122 (99%)	1e-68

Q9H2A9	N-acetylgalactosamine-4-O-sulfotrans ferase - Homo sapiens (Human), 424 aa.	62..183 303..424	120/122 (98%) 120/122 (98%)	4e-68
Q9BXH4	GalNAc-4-sulfotransferase 2 - Homo sapiens (Human), 443 aa.	62..179 325..442	77/118 (65%) 95/118 (80%)	1e-44
Q9BXH3	GalNAc-4-sulfotransferase 2 - Homo sapiens (Human), 358 aa.	62..179 240..357	77/118 (65%) 95/118 (80%)	1e-44
Q9BZW9	N-acetylgalactosamine 4-O-sulfotransferase 2 GalNAc4ST-2 - Homo sapiens (Human), 438 aa.	62..179 320..437	77/118 (65%) 95/118 (80%)	1e-44

Example 31.

The NOV31 clone was analyzed, and the nucleotide and encoded polypeptide
sequences are shown in Table 31A.

Table 31A. NOV31 Sequence Analysis

	SEQ ID NO: 133	2325 bp	
NOV31a, CG149008-01 DNA Sequence	CCCAGGCCGGACAAGCGTCCCGAAAGCCCCGGGAGAGACTAAGAAGCAATCCTCCCACGCGCTTTCT CCCACCCTCGGGCCACTGAGACGGAGGGACAGAGGGCCGCCCTCGCGCGGCCGAGGCCCCGCCTCCC GCTCGCCCGCCCGCGCCTCCAGCGGAAGCCGGAAGCAAAAGCGGGTCCTGCTAGCCCCGCGGCTCCG AACTCGGTGGTCCTGGAAGCTCCGCAGGATGGGGGAGAAGATGGCGGAAGAGGAGAGGTTCCCCAAT ACAACTCATGAGGGTTTCAATGTCACCCTCCACACCACCCTGGTTGTCACGACGAAACTGGTGCTCC CGACCCCTGGCAAGCCCATCCTCCCCGTGCAGACAGGGGAGCAGGCCCAGCAAGAGGAGCAGTCCAG CGGCATGACCATTTTCTTCAGCCTCCTTGTCCTAGCTATCTGCATCATATTGGTGCATTTACTGATC CGATACAGATTACATTTCTTGCCAGAGAGTGTTGCTGTTGTTTCTTTAGGTATTCTCATGGGAGCAG TTATAAAAATTATAGAGTTTAAAAAACTGGCGAATTGGAAGGAAGAAGAAATGTTTCGTCCAAACAT GTTTTTCCTCCTCCTGCTTCCCCCTATTATCTTTGAGTCTGGATATTCATTACACAAGGTGAGACTC AGGCACACATTGGGTAACTTCTTTCAAAATATTGGTTCCATCACCCTGTTTGCTGTTTTTGGGACGG CAATCTCCGCTTTTGTAGTAGGTGGAGGAATTTATTTTCTGGGTCAGGCTGATGTAATCTCTAAACT CAACATGACAGACAGTTTTGCGTTTGGCTCCCTAATATCTGCTGTCGATCCAGTGGCCACTATTGCC ATTTTCAATGCACTTCATGTGGACCCCGTGCTCAACATGCTGGTCTTTGGAGAAAGTATTCTCAACG ATGCAGTCTCCATTGTTCTGACCAACACAGCTGAAGGTTTAACAAGAAAAAATATGTCAGATGTCAG TGGGTGGCAAACATTTTTACAAGCCCTTGACTACTTCCTCAAAATGTTCTTTGGCTCTGCAGCGCTC GGCACTCTCACTGGCTTAATTTCTGCATTAGTGCTGAAGCATATTGACTTGAGGAAAACGCCTTCCT TGGAGTTTGGCATGATGATCATTTTTGCTTATCTGCCTTATGGGCTTGCAGAAGGAATCTCACTCTC AGGCATCATGGCCATCCTGTTCTCAGGCATCGTGATGTCCCACTACACGCACCATAACCTCTCCCCA GTCACCCAGATCCTCATGCAGCAGACCCTCCGCACCGTGGCCTTCTTATGTGAAACATGTGTGTTTG CATTTCTTGGCCTGTCCATTTTTAGTTTTCCTCACAAGTTTGAAATTTCCTTTGTCATCTGGTGCAT AGTGCTTGTACTATTTGGCAGAGCGGTAAACATTTTCCCTCTTTCCTACCTCCTGAATTTCTTCCGG GATCATAAAATCACACCGAAGATGATGTTCATCATGTGGTTTAGTGGCCTGCGGGGAGCCATCCCCT ATGCCCTGAGCCTACACCTGGAGCCCATGGAGAAGCGGCAGCTCATCGGCACCACCACCAT CGTCATCGTGCTCTTCACCATCCTGCTGCTGGGCGGCAGCACCATGCCCCCTCATTCGCCTCATGGAC ATCGAGGACGCCAAGGCACACCGCAGGAACAAGAAGGACGTCAACCTCAGCAAGACTGAGAAGATGG GCAACACTGTGGAGTCGGAGCACCTGTCGGAGCTCACGGAGGAGGAGTACGAGGCCCACTACATCAG GCGGCAGGACCTTAAGGGCTTCGTGTGGCTGGACGCCAAGTACCTGAACCCCTTCTTCACTCGGAGG CTGACGCAGGAGGACCTGCACCACGGGCGCATCCAGATGAAAACTCTCACCAACAAGTGGTACGAGG AGGTACGCCAGGGCCCCTCCGGCTCCGAGGACGACGAGCAGGAGCTGCTCTGACGCCAGGTGCCAAG GCTTCAGGCAGGCAGGCCCAGGATGGGCGTTTGCTGCGCACAGACACTCAGCAGGGGCCTCGCAGAG ATGCGTGCATCCAGCAGCCCCTTCAAGACATAAGAGGGCGGGGCGAGGTACTGGCTGCAGAGTCGCC TTAGTCCAGAACCTGACAGGCCTCTGGAGCCAGGCGACTTCTTGGGGAAACTGTCATCTCCCGACTCC TCCCTGAGCCAGCCTCCGCTCAGTGTGGCCTCTCAGCCCACAGAGGGGAGGGGAGCATGGGGCCAGGT GCCAGTCATCTGTGAAGCTAGGGCGCCTACCCCCCCCACCCGGAGGAC		
	ORF Start: ATG at 230		ORF Stop: TGA at 1994

	SEQ ID NO: 134	588 aa	MW at 66297.1kD
NOV31a, CG149008-01 Protein Sequence	MGEKMAEEERFPNTTHEGFNVTLHTTLVVTTKLVLPTPGKPILPVQTGEQAQQEEQSSGMTIFFSLL VLAICIILVHLLIRYRLHFLPESVAVVSLGILMGAVIKIIEFKKLANWKEEEMFRPNMFFLLLLPPI IFESGYSLHKVRLRHTLGNFFQNIGSITLFAVFGTAISAFVVGGGIYFLGQADVISKLNMTDSFAFG SLISAVDPVATIAIFNALHVDPVLNMLVFGESILNDAVSIVLTNTAEGLTRKNMSDVSGWQTFLQAL DYFLKMFFGSAALGTLTGLISALVLKHIDLRKTPSLEFGMMIIFAYLPYGLAEGISLSGIMAILFSG IVMSHYTHHNLSPVTQILMQQTLRTVAFLCETCVFAFLGLSIFSFPHKFEISFVIWCIVLVLFGRAV NIFPLSYLLNFFRDHKITPKMMFIMWFSGLRGAIPYALSLHLDLEPMEKRQLIGTTTIVIVLFTILL LGGSTMPLIRLMDIEDAKAHRRNKKDVNLSKTEKMGNTVESEHLSELTEEEYEAHYIRRQDLKGFVW LDAKYLNPFFTRRLTQEDLHHGRIQMKTLTNKWYEEVRQGPSGSEDDEQELL

Further analysis of the NOV31a protein yielded the following properties shown in Table 31B.

Table 31B. Protein Sequence Properties NOV31a	
PSort analysis:	0.8000 probability located in plasma membrane; 0.4000 probability located in Golgi body; 0.3000 probability located in endoplasmic reticulum (membrane); 0.3000 probability located in microbody (peroxisome)
SignalP analysis:	Cleavage site between residues 40 and 41

A search of the NOV31a protein against the Geneseq database, a proprietary database that contains sequences published in patents and patent publication, yielded several homologous proteins shown in Table 31C.

Table 31C. Geneseq Results for NOV31a				
Geneseq Identifier	Protein/Organism/Length [Patent #, Date]	NOV31a Residues/ Match Residues	Identities/ Similarities for the Matched Region	Expect Value
ABG61535	Human transporter and ion channel, TRICH5, Incyte ID 7476938CD1 - Homo sapiens, 671 aa. [WO200240541-A2, 23-MAY-2002]	1..588 91..671	581/588 (98%) 581/588 (98%)	0.0
AAM24062	Human EST encoded protein SEQ ID NO: 1587 - Homo sapiens, 315 aa. [WO200154477-A2, 02-AUG-2001]	274..588 1..315	315/315 (100%) 315/315 (100%)	0.0

AAB29621	Cat flea HMT Na/H transporter, SEQ ID NO:1868 - Ctenocephalides felis, 608 aa. [WO200061621-A2, 19-OCT-2000]	8..584 33..602	329/585 (56%) 416/585 (70%)	e-175
ABB59364	Drosophila melanogaster polypeptide SEQ ID NO 4884 - Drosophila melanogaster, 649 aa. [WO200171042-A2, 27-SEP-2001]	44..587 86..635	310/562 (55%) 399/562 (70%)	e-170
AAO14196	Human transporter and ion channel TRICH-13 - Homo sapiens, 631 aa. [WO200204520-A2, 17-JAN-2002]	117..547 125..542	166/439 (37%) 253/439 (56%)	2e-72

In a BLAST search of public sequence datbases, the NOV31a protein was found to have homology to the proteins shown in the BLASTP data in Table 31D.

5

Table 31D. Public BLASTP Results for NOV31a				
Protein Accession Number	Protein/Organism/Length	NOV31a Residues/ Match Residues	Identities/ Similarities for the Matched Portion	Expect Value
BAA76783	KIAA0939 protein - Homo sapiens (Human), 595 aa (fragment).	1..588 15..595	581/588 (98%) 581/588 (98%)	0.0
Q8R4D1	Na-H exchanger isoform NHE8 - Mus musculus (Mouse), 576 aa.	5..587 1..575	556/583 (95%) 565/583 (96%)	0.0
Q9Y507	DJ963K23.4 (Continues in dJ1041C10 (AL162615)) - Homo sapiens (Human), 437 aa (fragment).	152..588 1..437	437/437 (100%) 437/437 (100%)	0.0
Q9Y2E8	KIAA0939 protein - Homo sapiens (Human), 411 aa (fragment).	182..588 5..411	405/407 (99%) 406/407 (99%)	0.0
AAH34508	Hypothetical protein - Mus musculus (Mouse), 388 aa (fragment).	209..587 9..387	366/379 (96%) 374/379 (98%)	0.0

PFam analysis predicts that the NOV31a protein contains the domains shown in the Table 31E.

Table 31E. Domain Analysis of NOV31a			
Pfam Domain	NOV31a Match Region	Identities/ Similarities for the Matched Region	Expect Value
Na_H_Exchanger	62..485	141/465 (30%) 345/465 (74%)	3.1e-98

Example 32.

The NOV32 clone was analyzed, and the nucleotide and encoded polypeptide sequences are shown in Table 32A.

Table 32A. NOV32 Sequence Analysis			
	SEQ ID NO: 135	367 bp	
NOV32a, CG149350-01 DNA Sequence	ATGGCGGGGAGAAGGAAGCTCATCGCAGTGATCAGAGACAAGGACACGGTGACTGGTTTCCTGCTGG GCAGCATAGGGGAGCTTAACAAGAACTGCCACCCCAATTTCCTGGTGGTGGAGAAGGATACGACCAT CAATGAGATCGAAGACACTTTCCGGCAATTTCTAAACCGGGATGACACTGGCATCATCCTCATCAAC CAGTACATCGCAGAGATGGTGCAGCATGCCCTGGACACCCACCAGCACTCTATCCCTACTGTCCTGG AGATCCCCTCCAAGGAGCACCCATATGAGGACGCCAAGGACTCCACCCTGCGGAGGGCCAGGGGCAT GTTCACTGCCGAAGACCTGTGCTAGGGTCTTT		
	ORF Start: ATG at 1		ORF Stop: TAG at 358

	SEQ ID NO: 136	119 aa	MW at 13566.3kD
NOV32a, CG149350-01 Protein Sequence	MAGRRKLIAVIRDKDTVTGFLLGSIGELNKNCHPNFLVVEKDTTINEIEDTFRQFLNRDDTGIILIN QYIAEMVQHALDTHQHSIPTVLEIPSKEHPYEDAKDSTLRRARGMFTAEDLC		

	SEQ ID NO: 137	367 bp	
NOV32b, CG149350-02 DNA Sequence	ATGGCGGGGAGAAGGAAGCTCATCGCAGTGATCAGAGACAAGGACACGGTGACTGGTTTCCTGCTGG GCAGCATAGGGGAGCTTAACAAGAACTGCCACCCCAATTTCCTGGTGGTGGAGAAGGATACGACCAT CAATGAGATCGAAGACACTTTCCGGCAATTTCTAAACCGGGATGACACTGGCATCATCCTCATCAAC CAGTACATCGCAGAGATGGTGCAGCATGCCCTGGACACCCACCAGCACTCTATCCCTACTGTCCTGG AGATCCCCTCCAAGGAGCACCCATATGAGGACGCCAAGGACTCCACCCTGCGGAGGGCCAGGGGCAT GTTCACTGCCGAAGACCTGTGCTAGGGTCTTT		
	ORF Start: ATG at 1		ORF Stop: TAG at 358

	SEQ ID NO: 138	119 aa	MW at 13566.3kD
NOV32b, CG149350-02 Protein Sequence	MAGRRKLIAVIRDKDTVTGFLLGSIGELNKNCHPNFLVVEKDTTINEIEDTFRQFLNRDDTGIILIN QYIAEMVQHALDTHQHSIPTVLEIPSKEHPYEDAKDSTLRRARGMFTAEDLC		

Sequence comparison of the above protein sequences yields the following sequence relationships shown in Table 32B.

Table 32B. Comparison of NOV32a against NOV32b.		
Protein Sequence	**NOV32a Residues/ Match Residues**	**Identities/ Similarities for the Matched Region**
NOV32b	1..119 1..119	119/119 (100%) 119/119 (100%)

Further analysis of the NOV32a protein yielded the following properties shown in Table 32C.

Table 32C. Protein Sequence Properties NOV32a	
PSort analysis:	0.4852 probability located in mitochondrial matrix space; 0.4500 probability located in cytoplasm; 0.1957 probability located in mitochondrial inner membrane; 0.1957 probability located in mitochondrial intermembrane space
SignalP analysis:	No Known Signal Sequence Predicted

A search of the NOV32a protein against the Geneseq database, a proprietary database that contains sequences published in patents and patent publication, yielded several homologous proteins shown in Table 32D.

Table 32D. Geneseq Results for NOV32a				
Geneseq Identifier	**Protein/Organism/Length [Patent #, Date]**	**NOV32a Residues/ Match Residues**	**Identities/ Similarities for the Matched Region**	**Expect Value**

AAW27337	Human vacuolar ATPase 14 kDa subunit hV-14B - Homo sapiens, 119 aa. [JP09168390-A, 30-JUN-1997]	1..118 1..118	105/118 (88%) 108/118 (90%)	2e-54
AAW27336	Human vacuolar ATPase 14 kDa subunit hV-14A - Homo sapiens, 119 aa. [JP09168390-A, 30-JUN-1997]	1..118 1..118	104/118 (88%) 107/118 (90%)	8e-54
ABB62928	Drosophila melanogaster polypeptide SEQ ID NO 15576 - Drosophila melanogaster, 124 aa. [WO200171042-A2, 27-SEP-2001]	6..118 10..122	71/113 (62%) 91/113 (79%)	2e-38
ABB57798	Drosophila melanogaster polypeptide SEQ ID NO 186 - Drosophila melanogaster, 124 aa. [WO200171042-A2, 27-SEP-2001]	6..114 10..118	58/109 (53%) 84/109 (76%)	7e-29
AAG35989	Zea mays protein fragment SEQ ID NO: 44042 - Zea mays subsp. mays, 130 aa. [EP1033405-A2, 06-SEP-2000]	1..118 1..125	56/125 (44%) 85/125 (67%)	1e-27

In a BLAST search of public sequence datbases, the NOV32a protein was found to have homology to the proteins shown in the BLASTP data in Table 32E.

5

Table 32E. Public BLASTP Results for NOV32a				
Protein Accession Number	Protein/Organism/Length	NOV32a Residues/ Match Residues	Identities/ Similarities for the Matched Portion	Expect Value
P50408	Vacuolar ATP synthase subunit F (EC 3.6.3.14) (V-ATPase F subunit) (Vacuolar proton pump F subunit) (V-ATPase 14 kDa subunit) - Rattus norvegicus (Rat), 119 aa.	1..118 1..118	104/118 (88%) 108/118 (91%)	1e-53

Q16864	Vacuolar ATP synthase subunit F (EC 3.6.3.14) (V-ATPase F subunit) (Vacuolar proton pump F subunit) (V-ATPase 14 kDa subunit) - Homo sapiens (Human), 119 aa.	1..118 1..118	104/118 (88%) 107/118 (90%)	2e-53
Q9D1K2	1110004G16Rik protein - Mus musculus (Mouse), 119 aa.	1..118 1..118	103/118 (87%) 108/118 (91%)	5e-53
Q28029	Vacuolar ATP synthase subunit F (EC 3.6.3.14) (V-ATPase F subunit) (Vacuolar proton pump F subunit) (V-ATPase 14 kDa subunit) - Bos taurus (Bovine), 110 aa (fragment).	10..118 1..109	97/109 (88%) 100/109 (90%)	7e-50
Q9I8H3	Vacuolar ATP synthase subunit F (EC 3.6.3.14) (V-ATPase F subunit) (Vacuolar proton pump F subunit) (V-ATPase 14 kDa subunit) - Xenopus laevis (African clawed frog), 110 aa (fragment).	10..118 1..109	83/109 (76%) 94/109 (86%)	7e-43

PFam analysis predicts that the NOV32a protein contains the domains shown in the Table 32F.

5

Table 32F. Domain Analysis of NOV32a			
Pfam Domain	NOV32a Match Region	Identities/ Similarities for the Matched Region	Expect Value
ATP-synt_F	8..108	51/107 (48%) 90/107 (84%)	9.2e-43

Example 33.

10   The NOV33 clone was analyzed, and the nucleotide and encoded polypeptide sequences are shown in Table 33A.

Table 33A. NOV33 Sequence Analysis

SEQ ID NO: 139	1510 bp	
NOV33a, CG149463-01 DNA Sequence	ATGGGTTCAGACTTTATGCCCTGAAAAGATCCTTCCAGCCCTGGCCATCTTGGACTTCTGGAGCTAC CCTGGCTCACAGGGGTCTTGTTGCCCTGGGTGTCCCCAGTTCTTGAAAAGAATCAGCCTGGGAGGGG CCACACCCTGACCATCCCCCTTTATCCCTTCTGAGATGTTTGTTAGGAAGTCTGGGTCCAGGGGATA TCATTTCTTGTTCCATCCATGCAGGGGTTGCTTACCTCGGGTAGGAAACCCTCAGGCGGTGGCAGGT GCACAGGTAGGGGAGGATGGAGAGGGCAGTGGTGCCTGAAGCCCTGGATGGGCGGAGCTGACCCCCC AACACCAACTCTATCATGCCTGCTCCTCCCTGTCCCCCCAGAGCTGCCTGATCATTGCTACAGAATG AACTCTAGCCCAGCTGGGACCCCAAGTCCACAGCCCTCCAGGGCCAATGGGAACATCAACCTGGGGC CTTCAGCCAACCCAAATGCCCAGCCCACGGACTTCGACTTCCTCAAAGTCATCGGCAAAGGGAACTA CGGGAAGGTCCTACTGGCCAAGCGCAAGTCTGATGGGGCGTTCTATGCAGTGAAGGTACTACAGAAA AAGTCCATCTTAAAGAAGAAAGAGCAGAGCCACATCATGGCAGAGCGCAGTGTGCTTCTGAAGAACG TGCGGCACCCCTTCCTCGTGGGCCTGCGCTACTCCTTCCAGACACCTGAGAAGCTCTACTTCGTGCT CGACTATGTCAACGGGGGAGAGCTCTTCTTCCACCTGCAGCGGGAGCGCCGGTTCCTGGAGCCCCGG GCCAGGTTCTACGCTGCTGAGGTGGCCAGCGCCATTGGCTACCTGCACTCCCTCAACATCATTTACA GGGATCTGAAACCAGAGAACATTCTCTTGGACTGCCAGTACTTGGCACCTGAAGTGCTTCGGAAAGA GCCTTATGATCGAGCAGTGGACTGGTGGTGCCTTGGGGGCAGTCCTCTACGAGATGCTCCATGGCCTG CCGCCCTTCTACAGCCAAGATGTATCCCAGATGTATGAGAACATTCTGCACCAGCCGCTACAGATCC CCGGAGGCCGGACAGTGGCCGCCTGTGACCTCCTGCAAAGCCTTCTCCACAAGGACCAGAGGCAGCG GCTGGGCTCCAAAGCAGACTTTCTTGAGATTAAGAACCATGTATTCTTCAGCCCCATAAACTGGGAT GACCTGTACCACAAGAGGCTAACTCCACCCTTCAACCCAAATGTGACAGGACCTGCTGACTTGAAGC ATTTTGACCCAGAGTTCACCCAGGAAGCTGTGTCCAAGTCCATTGGCTGTACCCCCGACACTGTGGC CAGCAGCTCTGGGGCCTCAAGTGCATTCCTGGGATTTTCTTATGCGCCAGAGGATGATGACATCTTG GATTGTTAGAAGAGAAGGGCCTGTGAAACTACTGAGGCCAGCTGGTATTAGTAAGGAATTACCTTCA GCTGCTAGGAAGAGCGACTCAAACTAACAATGGCTT

ORF Start: ATG at 220		ORF Stop: TAG at 1414

SEQ ID NO: 140	398 aa	MW at 44552.5kD
NOV33a, CG149463-01 Protein Sequence	MQGLLTSGRKPSGGGRCTGRGGWRGQWCLKPWMGGADPPTPTLSCLLLPVPPELPDHCYRMNSSPAG TPSPQPSRANGNINLGPSANPNAQPTDFDFLKVIGKGNYGKVLLAKRKSDGAFYAVKVLQKKSILKK KEQSHIMAERSVLLKNVRHPFLVGLRYSFQTPEKLYFVLDYVNGGELFFHLQRERRFLEPRARFYAA EVASAIGYLHSLNIIYRDLKPENILLDCQYLAPEVLRKEPYDRAVDWWCLGAVLYEMLHGLPPFYSQ DVSQMYENILHQPLQIPGGRTVAACDLLQSLLHKDQRQRLGSKADFLEIKNHVFFSPINWDDLYHKR LTPPFNPNVTGPADLKHFDPEFTQEAVSKSIGCTPDTVASSSGASSAFLGFSYAPEDDDILDC

Further analysis of the NOV33a protein yielded the following properties shown in Table 33B.

Table 33B. Protein Sequence Properties NOV33a	
PSort analysis:	0.4500 probability located in cytoplasm; 0.2677 probability located in microbody (peroxisome); 0.1859 probability located in lysosome (lumen); 0.1000 probability located in mitochondrial matrix space
SignalP analysis:	No Known Signal Sequence Predicted

A search of the NOV33a protein against the Geneseq database, a proprietary database that contains sequences published in patents and patent publication, yielded several homologous proteins shown in Table 33C.

5

Table 33C. Geneseq Results for NOV33a				
Geneseq Identifier	Protein/Organism/Length [Patent #, Date]	NOV33a Residues/ Match Residues	Identities/ Similarities for the Matched Region	Expect Value
AAY95276	Human serum and glucocorticoid-induced protein kinase 2-beta - Homo sapiens, 427 aa. [WO200035946-A1, 22-JUN-2000]	1..398 1..427	398/427 (93%) 398/427 (93%)	0.0
AAM25594	Human protein sequence SEQ ID NO:1109 - Homo sapiens, 382 aa. [WO200153455-A2, 26-JUL-2001]	53..398 8..382	346/375 (92%) 346/375 (92%)	0.0
AAE22765	Human serum and glucocoticoid-induced protein kinase, SGK2-alpha - Homo sapiens, 367 aa. [WO200224947-A2, 28-MAR-2002]	61..398 1..367	338/367 (92%) 338/367 (92%)	0.0
AAB65708	Novel protein kinase, SEQ ID NO: 237 - Homo sapiens, 367 aa. [WO200073469-A2, 07-DEC-2000]	61..398 1..367	337/367 (91%) 338/367 (91%)	0.0
AAB65615	Novel protein kinase, SEQ ID NO: 141 - Mus musculus, 244 aa. [WO200073469-A2, 07-DEC-2000]	184..398 1..244	215/244 (88%) 215/244 (88%)	e-122

In a BLAST search of public sequence datbases, the NOV33a protein was found to have homology to the proteins shown in the BLASTP data in Table 33D.

10

Table 33D. Public BLASTP Results for NOV33a

Protein Accession Number	Protein/Organism/Length	NOV33a Residues/ Match Residues	Identities/ Similarities for the Matched Portion	Expect Value
Q9HBY8	Protein kinase – Homo sapiens (Human), 427 aa.	1..398 1..427	398/427 (93%) 398/427 (93%)	0.0
Q9UKG6	Protein kinase (DJ138B7.2) (Serum/glucocorticoid regulated kinase 2) (Similar to serum/glucocorticoid regulated kinase 2) – Homo sapiens (Human), 367 aa.	61..398 1..367	338/367 (92%) 338/367 (92%)	0.0
Q8R0P6	Serum/glucocorticoid regulated kinase 2 – Mus musculus (Mouse), 366 aa.	61..397 1..365	317/366 (86%) 326/366 (88%)	0.0
O73927	S-sgk2 - Squalus acanthias (Spiny dogfish), 594 aa.	70..396 236..594	235/359 (65%) 277/359 (76%)	e-133
O73926	S-sgk1 - Squalus acanthias (Spiny dogfish), 433 aa.	61..396 60..433	239/374 (63%) 282/374 (74%)	e-132

PFam analysis predicts that the NOV33a protein contains the domains shown in the Table 33E.

5

Table 33E. Domain Analysis of NOV33a			
Pfam Domain	NOV33a Match Region	Identities/ Similarities for the Matched Region	Expect Value
pkinase	95..228	54/135 (40%) 116/135 (86%)	5e-39
pkinase	231..323	35/128 (27%) 69/128 (54%)	1.5e-21
pkinase_C	324..393	25/73 (34%) 47/73 (64%)	3.1e-15

Example 34.

10 The NOV34 clone was analyzed, and the nucleotide and encoded polypeptide sequences are shown in Table 34A.

Table 34A. NOV34 Sequence Analysis

	SEQ ID NO: 141	2152 bp	
NOV34a, CG149536-01 DNA Sequence	GGGGGGCCTGAGCCTCTCCGCCGGCGCAGGCTCTGCTCGCGCCAGCTCGCTCCCGCAGCCATGCCCA CCACCATCGAGCGGGAGTTCGAAGAGTTGGATACTCAGCGTCGCTGGCAGCCGCTGTACTTGGAAAT TCGAAATGAGTCCCATGACTATCCTCATAGAGTGGCCAAGTTTCCAGAAAACAGAAATCGAAACAGA TACAGAGATGTAAGCCCATATGATCACAGTCGTGTTAAACTGCAAAATGCTGAGAATGATTATATTA ATGCCAGTTTAGTTGACATAGAAGAGGCACAAAGGAGTTACATCTTAACACAGGGACCACTTCCTAA CACATGCTGCCATTTCTGGCTTATGGTTTGGCAGCAGAAGACCAAAGCAGTTGTCATGCTGAACCGC ATTGTGGAGAGAGAATCGAGTGGTGAAACCAGAACAATATCTCACTTTCATTATACTACCTGGCCAG ATTTTGGAGTCCCTGAATCACCAGCTTCATTTCTCAATTTCTTGTTTAAAGTGAGAGAATCTGGCTC CTTGAACCCTGACCATGGGCCTGCGGTGATCCACTGTAGTGCAGGCATTGGGCGCTCTGGCACCTTC TCTCTGGTAGACACTTGTCTTGTTTTGATGGAAAAAGGAGATGATATTAACATAAAACAAGTGTTAC TGAACATGAGAAAATACCGAATGGGTCTTATTCAGACCCCAGATCAACTGAGATTCTCATACATGGC TATAATAGAAGGAGCAAAATGTATAAAGGGAGATTCTAGTATACAGAAACGATGGAAAGAACTTTCT AAGGAAGACTTATCTCCTGCCTTTGATCATTCACCAAACAAAATAATGACTGAAAAATACAATGGGA ACAGAATAGGTCTAGAAGAAGAAAACTGACAGGTGACCGATGTACAGGACTTTCCTCTAAAATGCA AGATACAATGGAGGAGAACAGTGAGAGTGCTCTACGGAAACGTATTCGAGAGGACAGAAAGGCCACC ACAGCTCAGAAGGTGCAGCAGATGAAACAGAGGCTAAATGAGAATGAACGAAAAAGAAAAAGGTGGT TATATTGGCAACCTATTCTCACTAAGATGGGGTTTATGTCAGTCATTTTGGTTGGCGCTTTTGTTGG CTGGAGACTGTTTTTTCAGCAAAATGCCCTATAAACAATTAATTTTGCCCAGCAAGCTTCTGCACTA GTAACTGACAGTGCTACATTAATCATAGGGGTTTGTCTGCAGCAAACGCCTCATATCCCAAAAACGG TGCAGTAGAATAGACATCAACCAGATAAGTGATATTTACAGTCACAAGCCCAACATCTCAGGACTCT TGACTGCAGGTTCCTCTGAACCCCAAACTGTAAATGGCTGTCTAAAATAAAGACATTCATGTTTGTT AAAAACTGGTAAATTTTGCAACCTGTATTCATACATGTCAAACACAGTATTTCACCTGACCAACATTG AGATATCCTTTATCACAGGATTTGTTTTTGGAGGCTATCTGGATTTTAACCTGCACTTGATATAAGC AATAAATATTGTGGTTTTATCTACGTTATTGGAAAGAAAATGACATTTAAATAATGTGTGTAATGTA TAATGTACTATTGACATGGCCATCAACACTTTTATTCTTAAGCATTTCAGGGTAAATATATTTTATA AGTATCTATTTAATCTTTTGTAGTTAACTGTACTTTTTAAGAGCTCAATTTGAAAAATCTGTTACTA AAAAAAAAATTGTATGTCGATTGAATTGTACTGGATACATTTTCCATTTTTCTAAAAGAAGTTTG ATATGAGCAGTTAGAAGTTGGAATAAGCAATTTCTACTATATATTGCATTTCTTTTATGTTTTACAG TTTTCCCCATTTTAAAAAGAAAAGCAAACAAAGAAACAAAAGTTTTTCCTAAAAATATCTTTGAAGG AAAATTCTCCTTACTGGGATAGTCAGGTAAACAGTTGGTCAAGACTTTGTAAAGAAATTGGTTTCTG TAAATCCCATTATTGATATGTTTATTTTTCATGAAAATTTCAATGTAGTTGGGGTAGATTATGATTT AGGAAGCAAAAGTAAGAAGCAGCATTTTATGATTCATAATTTCAGTTTACTAGACTGAAGTTTTGAA GTAAACCC
	ORF Start: ATG at 61		ORF Stop: TAA at 1171

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	SEQ ID NO: 142	370 aa	MW at 43248.9kD
NOV34a, CG149536-01 Protein Sequence	MPTTIEREFEELDTQRRWQPLYLEIRNESHDYPHRVAKFPENRNRNRYRDVSPYDHSRVKLQNAEND YINASLVDIEEAQRSYILTQGPLPNTCCHFWLMVWQQKTKAVVMLNRIVERESSGETRTISHFHYTT WPDFGVPESPASFLNFLFKVRESGSLNPDHGPAVIHCSAGIGRSGTFSLVDTCLVLMEKGDDINIKQ VLLNMRKYRMGLIQTPDQLRFSYMAIIEGAKCIKGDSSIQKRWKELSKEDLSPAFDHSPNKIMTEKY NGNRIGLEEEKLTGDRCTGLSSKMQDTMEENSESALRKRIREDRKATTAQKVQQMKQRLNENERKRK RWLYWQPILTKMGFMSVILVGAFVGWRLFFQQNAL

10   Further analysis of the NOV34a protein yielded the following properties shown in Table 34B.

Table 34B. Protein Sequence Properties NOV34a

PSort analysis:	0.8500 probability located in endoplasmic reticulum (membrane); 0.4400 probability located in plasma membrane; 0.3000 probability located in nucleus; 0.1000 probability located in mitochondrial inner membrane

SignalP analysis:	No Known Signal Sequence Predicted

A search of the NOV34a protein against the Geneseq database, a proprietary database that contains sequences published in patents and patent publication, yielded several homologous proteins shown in Table 34C.

Table 34C. Geneseq Results for NOV34a				
Geneseq Identifier	Protein/Organism/Length [Patent #, Date]	NOV34a Residues/ Match Residues	Identities/ Similarities for the Matched Region	Expect Value
AAR14114	Non-receptor linked protein tyrosine phosphatase - Homo sapiens, 415 aa. [WO9113989-A, 19-SEP-1991]	1..370 1..415	368/415 (88%) 369/415 (88%)	0.0
AAU91293	Human NOV8 protein - Homo sapiens, 415 aa. [WO200216600-A2, 28-FEB-2002]	1..370 1..415	337/415 (81%) 345/415 (82%)	0.0
ABP41882	Human ovarian antigen HOCPJ87, SEQ ID NO:3014 - Homo sapiens, 368 aa. [WO200200677-A1, 03-JAN-2002]	24..336 5..362	312/358 (87%) 313/358 (87%)	e-178
AAM25250	Human protein sequence SEQ ID NO:765 - Homo sapiens, 168 aa. [WO200153455-A2, 26-JUL-2001]	116..269 14..167	137/154 (88%) 145/154 (93%)	1e-77
AAB56662	Human prostate cancer antigen protein sequence SEQ ID NO:1240 - Homo sapiens, 180 aa. [WO200055174-A1, 21-SEP-2000]	1..124 29..152	123/124 (99%) 124/124 (99%)	1e-69

In a BLAST search of public sequence datbases, the NOV34a protein was found to have homology to the proteins shown in the BLASTP data in Table 34D.

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Table 34D. Public BLASTP Results for NOV34a				
Protein Accession Number	Protein/Organism/Length	NOV34a Residues/ Match Residues	Identities/ Similarities for the Matched Portion	Expect Value
P17706	Protein-tyrosine phosphatase, non-receptor type 2 (EC 3.1.3.48) (T- cell protein-tyrosine phosphatase) (TCPTP) - Homo sapiens (Human), 415 aa.	1..370 1..415	369/415 (88%) 370/415 (88%)	0.0
A33899	protein-tyrosine-phosphatase (EC 3.1.3.48), nonreceptor type 2 - human, 415 aa.	1..370 1..415	368/415 (88%) 369/415 (88%)	0.0
A60345	protein-tyrosine-phosphatase (EC 3.1.3.48) 11A - human, 387 aa.	1..336 1..381	334/381 (87%) 335/381 (87%)	0.0
Q922E7	Protein tyrosine phosphatase, non-receptor type 2 - Mus musculus (Mouse), 406 aa.	1..365 1..405	323/410 (78%) 338/410 (81%)	0.0
Q06180	Protein-tyrosine phosphatase, non-receptor type 2 (EC 3.1.3.48) (Protein-tyrosine phosphatase PTP-2) (MPTP) - Mus musculus (Mouse), 382 aa.	1..336 1..376	298/381 (78%) 312/381 (81%)	e-168

5   PFam analysis predicts that the NOV34a protein contains the domains shown in the Table 34E.

Table 34E. Domain Analysis of NOV34a			
Pfam Domain	NOV34a Match Region	Identities/ Similarities for the Matched Region	Expect Value
Y_phosphatase	42..229	99/272 (36%) 163/272 (60%)	5.5e-88

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Example 35.

The NOV35 clone was analyzed, and the nucleotide and encoded polypeptide sequences are shown in Table 35A.

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Table 35A. NOV35 Sequence Analysis			
	SEQ ID NO: 143	908 bp	
NOV35a, CG149964-01 DNA Sequence	CCCTTCTACCCAGAGGGTGAATGGGTATCTTTCCCGGAATAATCCTAATTTTTCTAAGGGTGAAGTT TGCAACGGCGGCCGTGACTGTAAGCGGACACCAGAAAAGTACCACTGTAAGTCATGAGATGTCTGGT CTGAATTGGAAACCCTTTGTATATGGCGGCCTTGCCTCTATCGTGGCTGAGTTTGGGACTTTCCCTG TGGACCTTACCAAAACACGACTTCAGGTTCAAGGCCAAAGCATTGATGCCCGTTTCAAAGAGATAAA ATATAGAGGGATGTTCCATGCGCTGTTTCGCATCTGTAAAGAGGAAGGTGTATTGGCTCTCTATTCA GGAATTGCTCCTGCGTTGCTAAGACAAGCATCATATGGCACCATTAAAATTGGGATTTACCAAAGCT TGAAGCGCTTATTCGTAGAACGTTTAGAAGATGAAACTCTTTTAATTAATATGATCTGTGGGGTAGT GTCAGGAGTGATATCTTCCACTATAGCCAATCCCACCGATGTTCTAAAGATTCGAATGCAGGCTCAA GGAAGCTTGTTCCAAGGGAGCATGATTGGAAGCTTTATCGATATATACCAACAAGAAGGCACCAGGG GTCTGTGGAGGGGTGTGGTTCCAACTGCTCAGCGTGCTGCCATCGTTGTAGGAGTAGAGCTACCAGT CTATGATATTACTAAGAAGCATTTAATATTGTCAGGAATGATGGGACATGTGGATCTCTATAAGGGC ACTGTTGATGGTATTTTAAAGATGTGGAAACATGAGGGCTTTTTTGCACTCTATAAAGGATTTTGGC CAAACTGGCTTCGGCTTGGACCCTGGAACATCATTTTTTTTTATTACATACGAGCAGGTAAAGAGGCT TCAAATCTAAGAACTGAATTATATGTGAGCCCAGCAC
	ORF Start: ATG at 21		ORF Stop: TAA at 879

	SEQ ID NO: 144	286 aa	MW at 32043.5kD
NOV35a, CG149964-01 Protein Sequence	MGIFPGIILIFLRVKFATAAVTVSGHQKSTTVSHEMSGLNWKPFVYGGLASIVAEFGTFPVDLTKTR LQVQGQSIDARFKEIKYRGMFHALFRICKEEGVLALYSGIAPALLRQASYGTIKIGIYQSLKRLFVE RLEDETLLINMICGVVSGVISSTIANPTDVLKIRMQAQGSLFQGSMIGSFIDIYQQEGTRGLWRGVV PTAQRAAIVVGVELPVYDITKKHLILSGMMGHVDLYKGTVDGILKMWKHEGFFALYKGFWPNWLRLG PWNIIFFITYEQVKRLQI		

	SEQ ID NO: 145		871 bp	
NOV35b, 309326356 DNA Sequence	CACCGGATCCACCATGGGTATCTTTCCCGGAATAATCCTAATTTTTCTAAGGGTGAAGTTTGCAACG GCGGCCGTGATTCACCAGAAAAGTACCACTGTAAGTCATGAGATGTCTGGTCTGAATTGGAAACCCT TTGTATATGGCGGCCTTGCCTCTATCGTGGCTGAGTTTGGGACTTTCCCTGTGGACCTTACCAAAAC ACGACTTCAGGTTCAAGGCCAAAGCATTGATGCCCGTTTCAAAGAGATAAAATATAGAGGGATGTTC CATGCGCTGTTTCGCATCTGTAAAGAGGAAGGTGTATTGGCTCTCTATTCAGGAATTGCTCCTGCGT TGCTAAGACAAGCATCATATGGCACCATTAAAATTGGGATTTACCAAAGCTTGAAGCGCTTATTCGT AGAACGTTTAGAAGATGAAACTCTTTTAATTAATATGATCTGTGGGGTAGTGTCAGGAGTGATATCT TCCACTATAGCCAATCCCACCGATGTTCTAAAGATTCGAATGCAGGCTCAAGGAAGCTTGTTCCAAG GGAGCATGATTGGAAGCTTTATCGATATATACCAACAAGAAGGCACCAGGGGTCTGTGGAGGGGTGT GGTTCCAACTGCTCAGCGTGCTGCCATCGTTGTAGGAGTAGAGCTACCAGTCTATGATATTACTAAG AAGCATTTAATATTGTCAGGAATGATGGGACATGTGGATCTCTATAAGGGCACTGTTGATGGTATTT TAAAGATGTGGAAACATGAGGGCTTTTTTGCACTCTATAAAGGATTTTGGCCAAACTGGCTTCGGCT TGGACCCTGGAACATCATTTTTTTTTATTACATACGAGCAGGTAAAGAGGCTTCAAATCGTCGACGGC
	ORF Start: at 2		ORF Stop: end of sequence	

	SEQ ID NO: 146	290 aa	MW at 32429.9kD
NOV35b, 309326356 Protein Sequence	TGSTMGIFPGIILIFLRVKFATAAVIHQKSTTVSHEMSGLNWKPFVYGGLASIVAEFGTFPVDLTKT RLQVQGQSIDARFKEIKYRGMFHALFRICKEEGVLALYSGIAPALLRQASYGTIKIGIYQSLKRLFV ERLEDETLLINMICGVVSGVISSTIANPTDVLKIRMQAQGSLFQGSMIGSFIDIYQQEGTRGLWRGV VPTAQRAAIVVGVELPVYDITKKHLILSGMMGHVDLYKGTVDGILKMWKHEGFFALYKGFWPNWLRL GPWNIIFFITYEQVKRLQIVDG		

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	SEQ ID NO: 147	811 bp	
NOV35c, 309326444 DNA Sequence	CACCGGATCCGCCGTGATTCACCAGAAAAGTACCACTGTAAGTCATGAGATGTCTGGTCTGAATTGG AAACCCTTTGTATATGGCGGCCTTGCCTCTATCGTGGCTGAGTTTGGGACTTTCCCTGTGGACCTTA CCAAAACACGACTTCAGGTTCAAGGCCAAAGCATTGATGCCCGTTTCAAAGAGATAAAATATAGAGG GATGTTCCATGCGCTGTTTCGCATCTGTAAAGAGGAAGGTGTATTGGCTCTCTATTCAGGAATTGCT CCTGCGTTGCTAAGACAAGCATCATATGGCACCATTAAAATTGGGATTTACCAAAGCTTGAAGCGCT TATTCGTAGAACGTTTAGAAGATGAAACTCTTTTAATTAATATGATCTGTGGGGTAGTGTCAGGAGT GATATCTTCCACTATAGCCAATCCCACCGATGTTCTAAAGATTCGAATGCAAGCTCAAGGAAGCTTG TTCCAAGGGAGCATGATTGGAAGCTTTATCGATATATACCAACAAGAAGGCACCAGGGGTCTGTGGA GGGGTGTGGTTCCAACTGCTCAGCGTGCTGCCATCGTTGTAGGAGTAGAGCTACCAGTCTATGATAT TACTAAGAAGCATTTAATATTGTCAGGAATGATGGGACATGTGGATCTCTATAAGGGCACTGTTGAT GGTATTTTAAAGATGTGGAAACATGAGGGCTTTTTTGCACTCTATAAAGGATTTTGGCCAAACTGGC TTCGGCTTGGACCCTGGAACATCATTTTTTTTATTACATACGAGCAGGTAAAGAGGCTTCAAATCGT CGACGGC		
	ORF Start: at 2		ORF Stop: end of sequence

10

	SEQ ID NO: 148	270 aa	MW at 30239.1kD
NOV35c, 309326444 Protein Sequence	TGSAVIHQKSTTVSHEMSGLNWKPFVYGGLASIVAEFGTFPVDLTKTRLQVQGQSIDARFKEIKYRG MFHALFRICKEEGVLALYSGIAPALLRQASYGTIKIGIYQSLKRLFVERLEDETLLINMICGVVSGV ISSTIANPTDVLKIRMQAQGSLFQGSMIGSFIDIYQQEGTRGLWRGVVPTAQRAAIVVGVELPVYDI TKKHLILSGMMGHVDLYKGTVDGILKMWKHEGFFALYKGFWPNWLRLGPWNIIFFITYEQVKRLQIV DG		

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	SEQ ID NO: 149	761 bp	
NOV35d, 309326473 DNA Sequence	CACCGGATCCCTGAATTGGAAACCCTTTGTATATGGCGGCCTTGCCTCTATCGTGGCTGAGTTTGGG ACTTTCCCTGTGGACCTTACCAAAACACGACTTCAGGTTCAAGGCCAAAGCATTGATGCCCGTTTCA AAGAGATAAAATATAGAGGGATGTTCCATGCGCTGTTTCGCATCTGTAAAGAGGAAGGTGTATTGGC TCTCTATTCAGGAATTGCTCCTGCGTTGCTAAGACAAGCATCATATGGCACCATTAAAATTGGGATT TACCAAAGCTTGAAGCGCTTATTCGTAGAACGTTTAGAAGATGAAACTCTTTTAATTAATATGATCT GTGGGGTAGTGTCAGGAGTGATATCTTCCACTATAGCCAATCCCACCGATGTTCTAAAGATTCGAAT GCAGGCTCAAGGAAGCTTGTTCCAAGGGAGCATGATTGGAAGCTTTATCGATATATACCAACAAGAA GGCACCAGGGGTCTGTGGAGGGGTGTGGTTCCAACTGCTCAGCGTGCTGCCATCGTTGTAGGAGTAG AGCTACCAGTCTATGATATTACTAAGAAGCATTTAATATTGTCAGGAATGATGGGACATGTGGATCT CTATAAGGGCACTGTTGATGGTATTTTAAAGATGTGGAAACATGAGGGCTTTTTTGCACTCTATAAA GGATTTTGGCCAAACTGGCTTCGGCTTGGACCCTGGAACATCATTTTTTTTATTACATACGAGCAGG TAAAGAGGCTTCAAATCGTCGACG		
	ORF Start: at 2		ORF Stop: end of sequence

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	SEQ ID NO: 150	254 aa	MW at 28488.2kD
NOV35d, 309326473 Protein Sequence	TGSLNWKPFVYGGLASIVAEFGTFPVDLTKTRLQVQGQSIDARFKEIKYRGMFHALFRICKEEGVLA LYSGIAPALLRQASYGTIKIGIYQSLKRLFVERLEDETLLINMICGVVSGVISSTIANPTDVLKIRM QAQGSLFQGSMIGSFIDIYQQEGTRGLWRGVVPTAQRAAIVVGVELPVYDITKKHLILSGMMGHVDL YKGTVDGILKMWKHEGFFALYKGFWPNWLRLGPWNIIFFITYEQVKRLQIVDX

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	SEQ ID NO: 151	1019 bp	
NOV35e, CG149964-02 DNA Sequence	CTACCCAGAGGGTGAATGGGTATCTTTCCCGGAATAATCCTAATTTTTCTAAGGGTGAAGTTTGCAA CGGCGGCCGTGACTGTAAGCGGACACCAGAAAAGTACCACTGTAAGTCATGAGATGTCTGGTCTGAA TTGGAAACCCTTTGTATATGGCGGCCTTGCCTCTATCGTGGCTGAGTTTGGGACTTTCCCTGTGGAC CTTACCAAAACACGACTTCAGGTTCAAGGCCAAAGCATTGATGCCCGTTTCAAAGAGATAAAATATA GAGGGATGTTCCATGCGCTGTTTCGCATCTGTAAAGAGGAAGGTGTATTGGCTCTCTATTCAGGAAT TGCTCCTGCGTTGCTAAGACAAGCATCATATGGCACCATTAAAATTGGGATTTACCAAAGCTTGAAG CGCTTATTCGTAGAACGTTTAGAAGATGAAACTCTTTTAATTAATATGATCTGTGGGGTAGTGTCAG GAGTGATATCTTCCACTATAGCCAATCCCACCGATGTTCTAAAGATTCGAATGCAGGCTCAAGGAAG CTTGTTCCAAGGGAGCATGATTGGAAGCTTTATCGATATATACCAGCAAGAAGGCACCAGGGGTCTG TGGAGGGGTGTGGTTCCAACTGCTCAGCGTGCTGCCATCGTTGTAGGAGTAGAGCTACCAGTCTATG ATATTACTAAGAAGCATTTAATATTGTCAGGAATGATGGGCGATACAATTTTAACTCACTTCGTTTC CAGCTTTACATGTGGTTTGGCTGGGGCTCTGGCCTCCAACCCGGTTGATGTGGTTCGAACTCGCATG ATGAACCAGAGGGCAATCGTGGGACATGTGGATCTCTATAAGGGCACTGTTGATGGTATTTTAAAGA TGTGGAAACATGAGGGCTTTTTTGCACTCTATAAAGGATTTTGGCCAAACTGGCTTCGGCTTGGACC CTGGAACATCATTTTTTTTATTACATACGAGCAGGTAAAGAGGCTTCAAATCTAAGAACTGAATTAT ATGTGAGCCCAGCC
	ORF Start: ATG at 16		ORF Stop: TAA at 991

10

	SEQ ID NO: 152	325 aa	MW at 36175.2kD
NOV35e, CG149964-02 Protein Sequence	MGIFPGIILIFLRVKFATAAVTVSGHQKSTTVSHEMSGLNWKPFVYGGLASIVAEFGTFPVDLTKTR LQVQGQSIDARFKEIKYRGMFHALFRICKEEGVLALYSGIAPALLRQASYGTIKIGIYQSLKRLFVE RLEDETLLINMICGVVSGVISSTIANPTDVLKIRMQAQGSLFQGSMIGSFIDIYQQEGTRGLWRGVV PTAQRAAIVVGVELPVYDITKKHLILSGMMGDTILTHFVSSFTCGLAGALASNPVDVVRTRMMNQRA IVGHVDLYKGTVDGILKMWKHEGFFALYKGFWPNWLRLGPWNIIFFITYEQVKRLQI

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Sequence comparison of the above protein sequences yields the following sequence relationships shown in Table 35B.

Table 35B. Comparison of NOV35a against NOV35b through NOV35e.		
Protein Sequence	**NOV35a Residues/ Match Residues**	**Identities/ Similarities for the Matched Region**
NOV35b	1..286 5..287	282/286 (98%) 282/286 (98%)
NOV35c	26..286 7..267	261/261 (100%) 261/261 (100%)

NOV35d	39..286	248/248 (100%)
	4..251	248/248 (100%)
NOV35e	1..286	286/325 (88%)
	1..325	286/325 (88%)

Further analysis of the NOV35a protein yielded the following properties shown in Table 35C.

Table 35C. Protein Sequence Properties NOV35a	
PSort analysis:	0.4600 probability located in plasma membrane; 0.2648 probability located in microbody (peroxisome); 0.1000 probability located in endoplasmic reticulum (membrane); 0.1000 probability located in endoplasmic reticulum (lumen)
SignalP analysis:	Cleavage site between residues 20 and 21

A search of the NOV35a protein against the Geneseq database, a proprietary database that contains sequences published in patents and patent publication, yielded several homologous proteins shown in Table 35D.

Table 35D. Geneseq Results for NOV35a				
Geneseq Identifier	Protein/Organism/Length [Patent #, Date]	NOV35a Residues/ Match Residues	Identities/ Similarities for the Matched Region	Expect Value
AAY94665	Human uncoupling protein (UCP5) amino acid sequence - Homo sapiens, 325 aa. [WO200032624-A2, 08-JUN-2000]	1..286 1..325	284/325 (87%) 285/325 (87%)	e-158
ABG33878	Human secreted protein encoded by gene 16 - Homo sapiens, 334 aa. [WO200226931-A2, 04-APR-2002]	1..286 1..334	284/334 (85%) 285/334 (85%)	e-155
AAE06056	Human gene 16 encoded secreted protein HMIAP86, SEQ ID NO:118 - Homo sapiens, 334 aa. [WO200151504-A1, 19-JUL-2001]	1..286 1..334	284/334 (85%) 285/334 (85%)	e-155

| AAY87079 | Human secreted protein sequence SEQ ID NO:118 - Homo sapiens, 335 aa. [WO200004140-A1, 27-JAN-2000] | 1..286 1..334 | 284/334 (85%) 285/334 (85%) | e-155 |
| AAY94666 | Human uncoupling protein isoform hUCP5S amino acid sequence - Homo sapiens, 322 aa. [WO200032624-A2, 08-JUN-2000] | 1..286 1..322 | 281/325 (86%) 282/325 (86%) | e-154 |

In a BLAST search of public sequence datbases, the NOV35a protein was found to have homology to the proteins shown in the BLASTP data in Table 35E.

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Table 35E. Public BLASTP Results for NOV35a				
Protein Accession Number	Protein/Organism/Length	NOV35a Residues/ Match Residues	Identities/ Similarities for the Matched Portion	Expect Value
O95258	Brain mitochondrial carrier protein-1 (BMCP-1) (Mitochondrial uncoupling protein 5) (UCP 5) (Solute carrier family 25, member 14) - Homo sapiens (Human), 325 aa.	1..286 1..325	284/325 (87%) 285/325 (87%)	e-157
Q9Z2B2	Brain mitochondrial carrier protein-1 (BMCP-1) (Mitochondrial uncoupling protein 5) (UCP 5) (Solute carrier family 25, member 14) - Mus musculus (Mouse), 325 aa.	1..286 1..325	276/325 (84%) 283/325 (86%)	e-154
Q9EP88	Brain mitochondrial carrier protein BMCP1 (Brain mitochondrial carrier protein-1) - Rattus norvegicus (Rat), 325 aa.	1..286 1..325	274/325 (84%) 282/325 (86%)	e-153
Q9JMH0	Brain mitochondrial carrier protein-1 - Rattus norvegicus (Rat), 322 aa.	1..286 1..322	271/325 (83%) 279/325 (85%)	e-149
Q8R206	Similar to RIKEN cDNA 4933433D23 gene - Mus musculus (Mouse), 210 aa.	36..232 1..197	160/197 (81%) 176/197 (89%)	1e-87

PFam analysis predicts that the NOV35a protein contains the domains shown in the Table 35F.

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Table 35F. Domain Analysis of NOV35a			
Pfam Domain	NOV35a Match Region	Identities/ Similarities for the Matched Region	Expect Value
mito_carr	39..138	39/126 (31%) 78/126 (62%)	5.7e-24
mito_carr	140..231	29/125 (23%) 76/125 (61%)	4.4e-27
mito_carr	233..286	24/125 (19%) 46/125 (37%)	0.0072

Example 36.

The NOV36 clone was analyzed, and the nucleotide and encoded polypeptide

10   sequences are shown in Table 36A.

Table 36A. NOV36 Sequence Analysis			
	SEQ ID NO: 153	1144 bp	
NOV36a, CG150306-01 DNA Sequence	CGCGGGGCGCGCGGCGCGGGGCGGCCTGGCCGGCGGCGGCGGCGGCATGAAGGTCACGTCGCTCGAC GGGCGCCAGCTGCGCAAGATGCTCCGCAAGGAGGCGGCGGCGCGCTGCGTGGTGCTCGACTGCCGGC CCTATCTGGCCTTCGCTGCCTCGAACGTGCGCGGCTCGCTCAACGTCAACCTCAACTCGGTGGTGCT GGACCAGGGCAGCCGCCACTGGCAGAAGCTGCGAGAGGAGAGCGCCGCGCGTGTCGTCCTCACCTCG CTACTCGCTTGCCTACCCGCCGGCCCGCGGGTCTACTTCCTCAAAGGGGGATATGAGACTTTCTACT CGGAATATCCTGAGTGTTGCGTGGATGTAAAACCCATTTCACAAGAGAAGATTGAGAGTGAGAGAGC CCTCATCAGCCAGTGTGGAAAACCAGTGGTAAATGTCAGCTACAGGCCAGCTTATGACCAGGGTGGC CCAGTTGAAATCCTTCCCTTCCTCTACCTTGGAAGTGCCTACCATGCATCCAAGTGCGAGTTCCTCG CCAACTTGCACATCACAGCCCTGCTGAATGTCTCCCGACGGACCTCCGAGGCCTGCATGACCCACCT ACACTACAAATGGATCCCTGTGGAAGACAGCCACACCGGCTGACATTAGCTCCCACTTTCAAGAAGCA ATAGACTTCATTGACTGTGTCAGGGAAAAGGGAGGCAAGGTCCTGGTCCACTGTGAGGCTGGGATCT CCCGTTCACCCACCATCTGCATGGCTTACCTTATGAAGACCAAGCAGTTCCGCCTGAAGGAGGCCTT CGATTACATCAAGCAGAGGAGGAGCATGGTCTCGCCCAACTTTGGCTTCATGGGCCAGCTCCTGCAG TACGAATCTGAGATCCTGCCCTCCACGCCCAACCCCCAGCCTCCCTCCTGCCAAGGGGAGGCAGCAG GCTCTTCACTGATAGGCCATTTGCAGACACTGAGCCCTGACATGCAGGGTGCCTACTGCACATTCCC TGCCTCGGTGCTGGCACCGGTGCCTACCCACTCAACAGTCTCAGAGCTCAGCAGAAGCCCTGTGGCA ACGGCCACATCCTGCTAAAACTGGGATGGAGGAATCGGCCCAGCCCCAAGAGCAACTGTGATTTTTG TTTTT		
	ORF Start: ATG at 47		ORF Stop: TAA at 1088

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	SEQ ID NO: 154	347 aa	MW at 38362.6kD
NOV36a,	MKVTSLDGRQLRKMLRKEAAARCVVLDCRPYLAFAASNVRGSLNVNLNSVVLDQGSRHWQKLREESA		

| CG150306-01 Protein Sequence | ARVVLTSLLACLPAGPRVYFLKGGYETFYSEYPECCVDVKPISQEKIESERALTSQCGKPVVNVSYR PAYDQGGPVEILPFLYLGSAYHASKCEFLANLHITALLNVSRRTSEACMTHLHYKWIPVEDSHTADI SSHFQEAIDFIDCVREKGGKVLVHCEAGISRSPTICMAYLMKTKQFRLKEAFDYIKQRRSMVSPNFG FMGQLLQYESEILPSTPNPQPPSCQGEAAGSSLIGHLQTLSPDMQGAYCTFPASVLAPVPTHSTVSE LSRSPVATATSC |

Further analysis of the NOV36a protein yielded the following properties shown in Table 36B.

Table 36B. Protein Sequence Properties NOV36a	
PSort analysis:	0.4811 probability located in mitochondrial matrix space; 0.4500 probability located in cytoplasm; 0.1892 probability located in mitochondrial inner membrane; 0.1892 probability located in mitochondrial intermembrane space
SignalP analysis:	No Known Signal Sequence Predicted

A search of the NOV36a protein against the Geneseq database, a proprietary database that contains sequences published in patents and patent publication, yielded several homologous proteins shown in Table 36C.

Table 36C. Geneseq Results for NOV36a				
Geneseq Identifier	Protein/Organism/Length [Patent #, Date]	NOV36a Residues/ Match Residues	Identities/ Similarities for the Matched Region	Expect Value
ABB07842	Amino acid sequence of protein identified by Swissprot Accn No. Q16690 - Homo sapiens, 384 aa. [WO200220732-A2, 14-MAR-2002]	1..347 1..384	347/384 (90%) 347/384 (90%)	0.0
AAB66440	Human MAP-kinase phosphatase MKP-5 - Homo sapiens, 171 aa. [WO200102582-A1, 11-JAN-2001]	116..286 1..171	171/171 (100%) 171/171 (100%)	1e-97
AAE06784	Human dual-specificity phosphatase (DSP) protein, MKP-5 - Homo sapiens, 171 aa. [WO200157221-A2, 09-AUG-2001]	116..286 1..171	171/171 (100%) 171/171 (100%)	1e-97

| AAR63602 | MAP-kinase-phosphatase CL100 - Homo sapiens, 367 aa. [WO9423039-A, 13-OCT-1994] | 1..347 3..367 | 168/388 (43%) 220/388 (56%) | 5e-72 |
| AAU84270 | Human endometrial cancer related protein, DUSP1 - Homo sapiens, 367 aa. [WO200209573-A2, 07-FEB-2002] | 1..347 3..367 | 167/388 (43%) 219/388 (56%) | 1e-70 |

In a BLAST search of public sequence datbases, the NOV36a protein was found to have homology to the proteins shown in the BLASTP data in Table 36D.

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Table 36D. Public BLASTP Results for NOV36a				
Protein Accession Number	Protein/Organism/Length	NOV36a Residues/ Match Residues	Identities/ Similarities for the Matched Portion	Expect Value
Q16690	Dual specificity protein phosphatase 5 (EC 3.1.3.48) (EC 3.1.3.16) (Dual specificity protein phosphatase hVH3) - Homo sapiens (Human), 384 aa.	1..347 1..384	347/384 (90%) 347/384 (90%)	0.0
O54838	Dual specificity protein phosphatase 5 (EC 3.1.3.48) (EC 3.1.3.16) (MAP-kinase phosphatase CPG21) - Rattus norvegicus (Rat), 384 aa.	1..347 1..384	320/384 (83%) 336/384 (87%)	0.0
Q90W58	MAP kinase phosphatase XCL100(beta) protein - Xenopus laevis (African clawed frog), 369 aa.	13..347 15..369	164/378 (43%) 217/378 (57%)	9e-72
P28562	Dual specificity protein phosphatase 1 (EC 3.1.3.48) (EC 3.1.3.16) (MAP kinase phosphatase-1) (MKP-1) (Protein-tyrosine phosphatase CL100) (Dual specificity protein phosphatase hVH1) - Homo sapiens (Human), 367 aa.	1..347 3..367	167/388 (43%) 219/388 (56%)	3e-70

| O42253 | Dual specificity protein phosphatase 1 (EC 3.1.3.48) (EC 3.1.3.16) (MAP kinase phosphatase-1) (MPK-1) (MAP kinase phosphatase-1) - Gallus gallus (Chicken), 353 aa (fragment). | 15..344 4..353 | 166/366 (45%) 213/366 (57%) | 1e-68 |

PFam analysis predicts that the NOV36a protein contains the domains shown in the Table 36E.

Table 36E. Domain Analysis of NOV36a			
Pfam Domain	**NOV36a Match Region**	**Identities/ Similarities for the Matched Region**	**Expect Value**
Rhodanese	7..98	23/134 (17%) 66/134 (49%)	0.0052
DSPc	141..279	76/172 (44%) 132/172 (77%)	1.8e-70
Y_phosphatase	44..279	39/336 (12%) 144/336 (43%)	0.54

Example 37.

The NOV37 clone was analyzed, and the nucleotide and encoded polypeptide sequences are shown in Table 37A.

Table 37A. NOV37 Sequence Analysis			
	SEQ ID NO: 155	2277 bp	
NOV37a, CG150510-01 DNA Sequence	CGCGTTGTGGGCTCCCGCCGGGGTCCCCCGCGGCTGTCGCCGCCGCCTACGCCGCTGCCTCCGCCTT CCTGCCCCGCGTCGGGCCGGGCGCCACCTCCCCCCTGCCTCCCTCTCCGCTGTGGTCATTTAGGAAA TCGTAAATCATGTGAAGATGGGACTCTTGGTATTTGTGCGCAATCTGCTGCTAGCCCTCTGCCTCTT TCTGGTACTGGGATTTTTGTATTATTCTGCGTGGAAGCTACACTTACTCCAGTGGGAGGAGGACTCC AGTAAGTATAGTCACTCTAGCTCACCCCAGGAGAAGCCTGTTGCAGATTCAGTGGTTCTTTCCTTTG ACTCCGCTGGACAAACACTAGGCTCAGAGTATGATCGGTTGGGCTTCCTCCTGAATCTGGACTCTAA ACTGCCTGCTGAATTAGCCACCAAGTACGCAAACTTTTCAGAGGGAGCTTGCAAGCCTGGCTATGCT TCAGCCTTGATGACGGCCATCTTCCCCCGGTTCTCCAAGCCAGCACCCATGTTCCTGGATGACTCCT TTCGCAAGTGGGCTAGAATCCGGGAGTTCGTGCCGCCTTTTGGGATCAAAGGTCAAGACAATCTGAT CAAAGCCATCTTGTCAGTCACCAAAGAGTACCGCCTGACCCCTGCCTTGGACAGCCTCCGCTGCCGC CGCTGCATCATCGTGGGCAATGGAGGCGTTCTTGCCAACAAGTCTCTGGGGTCACGAATTGACGACT ATGACATTGTGGTGAGACTGAATTCAGCACCAGTGAAAGGCTTTGAGAAGGACGTGGGCAGCAAAAC GACACTGCGCATCACCTACCCCGAGGGCGCCATGCAGCGGCCTGAGCAGTACGAGCGCGATTCTCTC TTTGTCCTCGCCGGCTTCAAGTGGCAGGACTTTAAGTGGTTGAAATACATCGTCTACAAGGAGAGAG TGAGTGCATCGGATGGCTTCTGGAAATCTGTGGCCACTCGAGTGCCCAAGGAGCCCCCTGAGATTCG AATCCTCAACCCATATTTCATCCAGGAGGCCGCCTTCACCCTCATTGGCCTGCCCTTCAACAATGGC

	CTCATGGGCCGGGGGAACATCCCTACCCTTGGCAGTGTGGGCAGTGAGGATGGGCAGTACACGGGCTGTG
	ACGAGGTGGCAGTCGCAGGATTTGGCTATGACATGAGCACACCCAACGCACCCCTGCACTACTATGA
	GACCGTTCGCATGGCAGCCATCAAAGAGTCCTGGACGCACAATATCCAGCGAGAGAAAGAGTTTCTG
	CGGAAGCTGGTGAAAGCTCGCGTCATCACTGATCTAAGCAGTGGCATCTGAGTGGGCCCAGCACATG
	GCCATAGAGGCCCAGGCACCACCAGGAGCAGCAGCCAGCACCACCTACACAGGAGTCTTCAGACCCA
	GAGAAGGACGGTGCCAAGGGCCCCAGGGGCAGCAAGGCCTTGGTGGAGCAGCCAGAGCTGTGCCTGC
	TCAGCAGCCAGTCTCAGAGACCAGCCACTCAGCCTCATTCAGCATGGGTCCTTGATGCCAGAGGGCCA
	GCAGGCTCCTGGCTGTGCCCAGCAGGCCCAGCATGCAGGTGGTGGGACACTGGGCAGCAAGGCTGCT
	GCCGGAATCACTTCTCCAATCAGTGTTTGGTGTATTATCATTTTGTGAATTTGGGTAGGGGGGAGGG
	TAGGGATAATTTATTTTTAAATAAGGTTGGAGATGTCAAGTTGGGTTCACTTGCCATGCAGGAAGAG
	GCCCACTAGAGGGCCCATCAGGCAGTGTTACCTGTTAGCTCCCTGTGGGGCAGGAGTGCCAGGACCA
	GCCTGTACCTTGCTGTGGGGCTACAGGATGGTGGGCAGGATCTCAAGCCAGCCCCCTCCAGCTCATG
	ACACTGTTTGGCCTTTCTTGGGGAGAAGGCGGGGTATTCCCACTCACCAGCCCTAGCTGTCCCATGG
	GGAAACCCTGGAGCCATCCCTTCGGAGCCAACAAGACCGCCCCAGGGCTATAGCAGAAAGAACTTTA
	AAGCTCAGGAGGGTGACGCCCAGCTCCGCCTGCTGGGAAGAGCTCCCCTCCACAGCTGCAGCTGATC
	CATAGGACTACCGCAGGCCCGGACTCACCAACTTGCCACATGTTCTAGGTTTCAGCAACAAGACTGC
	CAGGTGGTTGGGTTCTGCCTTTAGCCTGGACCAAAGGGAAGTGAGGCCCAAGGAGCTTACCCAAGCT
	GTGGCAGCCGTCCCAGGCCACCCCCATGGAAGCAATAAAGCTCTTCCCTGTAAAAAAAAAAAAAAAA
ORF Start: ATG at 152		ORF Stop: TGA at 1322

| | SEQ ID NO: 156 | 390 aa | MW at 43785.1kD |
| NOV37a, CG150510-01 Protein Sequence | MGLLVFVRNLLLALCLFLVLGFLYYSAWKLHLLQWEEDSSKYSHSSSPQEKPVADSVVLSFDSAGQT LGSEYDRLGFLLNLDSKLPAELATKYANFSEGACKPGYASALMTAIFPRFSKPAPMFLDDSFRKWAR IREFVPPFGIKGQDNLIKAILSVTKEYRLTPALDSLRCRRCIIVGNGGVLANKSLGSRIDDYDIVVR LNSAPVKGFEKDVGSKTTLRITYPEGAMQRPEQYERDSLFVLAGFKWQDFKWLKYIVYKERVSASDG FWKSVATRVPKEPPEIRILNPYFIQEAAFTLIGLPFNNGLMGRGNIPTLGSVAVTMALHGCDEVAVA GFGYDMSTPNAPLHYYETVRMAAIKESWTHNIQREKEFLRKLVKARVITDLSSGI | | |

Further analysis of the NOV37a protein yielded the following properties shown in Table 37B.

Table 37B. Protein Sequence Properties NOV37a	
PSort analysis:	0.8200 probability located in outside; 0.2360 probability located in microbody (peroxisome); 0.1900 probability located in lysosome (lumen); 0.1000 probability located in endoplasmic reticulum (membrane)
SignalP analysis:	Cleavage site between residues 22 and 23

A search of the NOV37a protein against the Geneseq database, a proprietary database that contains sequences published in patents and patent publication, yielded several homologous proteins shown in Table 37C.

| Table 37C. Geneseq Results for NOV37a |

Geneseq Identifier	Protein/Organism/Length [Patent #, Date]	NOV37a Residues/ Match Residues	Identities/ Similarities for the Matched Region	Expect Value
AAY39960	Human alpha2-3 sialate transferase protein sequence - Homo sapiens, 375 aa. [JP11253163-A, 21-SEP-1999]	1..390 1..375	374/390 (95%) 375/390 (95%)	0.0
AAR65242	Human ST3N sialyltransferase - Homo sapiens, 375 aa. [WO9504816-A, 16-FEB-1995]	1..390 1..375	374/390 (95%) 375/390 (95%)	0.0
AAR63217	Human alpha-2,3-sialyltransferase (WM16) - Homo sapiens (melanoma WM266-4 cells), 375 aa. [WO9423021-A, 13-OCT-1994]	1..390 1..375	374/390 (95%) 375/390 (95%)	0.0
AAR62808	Alpha 2, 3-sialyl transferase - Homo sapiens, 375 aa. [JP06277052-A, 04-OCT-1994]	1..390 1..375	374/390 (95%) 375/390 (95%)	0.0
AAR41671	Rat sialyltransferase - Rattus rattus, 374 aa. [WO9318157-A, 16-SEP-1993]	1..390 1..374	361/390 (92%) 370/390 (94%)	0.0

In a BLAST search of public sequence datbases, the NOV37a protein was found to have homology to the proteins shown in the BLASTP data in Table 37D.

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Table 37D. Public BLASTP Results for NOV37a				
Protein Accession Number	Protein/Organism/Length	NOV37a Residues/ Match Residues	Identities/ Similarities for the Matched Portion	Expect Value
Q11203	CMP-N-acetylneuraminate-beta-1,4-gal actoside alpha-2,3- sialyltransferase (EC 2.4.99.6) (N-acetyllactosaminide alpha-2,3- sialyltransferase) (Gal beta-1,3(4) GlcNAc alpha-2,3 sialyltransferase) (ST3N) (Sialyltransferase 6) - Homo sapiens (Human), 375 aa.	1..390 1..375	374/390 (95%) 375/390 (95%)	0.0

Q922X5	Sialyltransferase (N-acetyllacosaminide alpha 2,3-sialyltransferase) - Mus musculus (Mouse), 374 aa.	1..390 1..374	361/390 (92%) 371/390 (94%)	0.0
Q9DBB6	Sialyltransferase (N-acetyllacosaminide alpha 2,3- sialyltransferase) - Mus musculus (Mouse), 374 aa.	1..390 1..374	360/390 (92%) 371/390 (94%)	0.0
Q02734	CMP-N-acetylneuraminate-beta-1,4-gal actoside alpha-2,3- sialyltransferase (EC 2.4.99.6) (N-acetyllactosaminide alpha-2,3- sialyltransferase) (Gal beta-1,3(4) GlcNAc alpha-2,3 sialyltransferase) (ST3N) (Sialyltransferase 6) - Rattus norvegicus (Rat), 374 aa.	1..390 1..374	361/390 (92%) 370/390 (94%)	0.0
P97325	CMP-N-acetylneuraminate-beta-1,4-gal actoside alpha-2,3- sialyltransferase (EC 2.4.99.6) (N-acetyllactosaminide alpha-2,3- sialyltransferase) (Gal beta-1,3(4) GlcNAc alpha-2,3 sialyltransferase) (ST3N) (Sialyltransferase 6) - Mus musculus (Mouse), 374 aa.	1..390 1..374	359/390 (92%) 370/390 (94%)	0.0

PFam analysis predicts that the NOV37a protein contains the domains shown in the Table 37E.

Table 37E. Domain Analysis of NOV37a			
Pfam Domain	NOV37a Match Region	Identities/ Similarities for the Matched Region	Expect Value
Glyco_transf_29	101..389	108/324 (33%) 270/324 (83%)	3.2e-116

Example 38.

The NOV38 clone was analyzed, and the nucleotide and encoded polypeptide sequences are shown in Table 38A.

Table 38A. NOV38 Sequence Analysis			
	SEQ ID NO: 157	1076 bp	
NOV38a,	CCCTTATGAAGACGGGACATTTTGAAATAGTCACCATGCTGCTGGCAACCATGATTCTAGTGGACAT		

236

CG150704-01 DNA Sequence	TTTCCAGGTGAAGGCTGAAGTGTTAGACATGGCAGATAATGGATTTCATCATGAATACCTGAAATGT ACGGACAGGATGGAAATTAAATACGTTCCCCAACTGCTAAAGGAGGAAAAAGCAAGCCACCAGCAAT TAGATACTGTGTGGGAAAATGCAAAAGCCAAATGGGCAGCCCGAAAGACTCAAATCTTTCTCCCTAT GAATTTTAAGGATAACCATGGAATAGCCCTGATGGCATATATTTCCGAAGCTCAAGAGCAAACTCCC TTTTACCATCTGTTCAGTGAAGCTGTGAAGATGGCTGGCCAATCTCGAGAAGATTATATCTATGGCT TCCAGTTCAAAGCTTTCCACTTTTACCTCACAAGAGCCCTGCAGTTGCTGAGAAAACCTTGTGAGGC CAGTTCCAAAACTGTGGTATATAGAACAAGCCAGGGCACTTCATTTACATTTGGAGGGCTAAACCAA GCCAGGTTTGGCCATTTTACCTTGGCATATTCAGCCAACCTCAGGCTGCTAATGACCAGCTCACTG TGTTATCCATCTACACATGCCTTGGAGTTGACATTGAAAATTTTCTTGATAAAGAAAGTGAAAGAAT TACTTTAATACCTCTGAATGAGGTTTTTCAAGTGTCACAGGAGGGGGCTGGCAATAACCTTATCCTT CAAAGCATAAACAAGACCTGCAGCCATTATGAGTGTGCATTTCTAGGTGGACTAAAAACCGAAAACT GTATTGAGAACCTAGAATATTTTCAACCCATCTATGTCTACAACCCTGGTGAGAAAAACCAGAAGCT TGAAGACCATAGTGAGAAAAACTGGAAGCTTGAAGACCATGGTGAGAAAAACCAGAAGCTTGAAGAC CATGCTCCAGGTCCAGTTCCTGTTCCAGGTCCCAAAAGCCATCCTTCTGCATCCTCGGGCAAACTGC TGCTTCCACAGTTTGGGATGGTCATCATTTTAATCAGTGTTTCTGCTATAAATCTCTTTGTTGCTCT GTAG
	ORF Start: ATG at 6		ORF Stop: TAG at 1074

	SEQ ID NO: 158	356 aa	MW at 40311.7kD
NOV38a, CG150704-01 Protein Sequence	MKTGHFEIVTMLLATMILVDIFQVKAEVLDMADNAFDDEYLKCTDRMEIKYVPQLLKEEKASHQQLD TVWENAKAKWAARKTQIFLPMNFKDNHGIALMAYISEAQEQTPFYHLFSEAVKMAGQSREDYIYGFQ FKAFHFYLTRALQLLRKPCEASSKTVVYRTSQGTSFTFGGLNQARFGHFTLAYSAKPQAANDQLTVL SIYTCLGVDIENFLDKESERITLIPLNEVFQVSQEGAGNNLILQSINKTCSHYECAFLGGLKTENCI ENLEYFQPIYVYNPGEKNQKLEDHSEKNWKLEDHGEKNQKLEDHAPGPVPVPGPKSHPSASSGKLLL PQFGMVIILISVSAINLFVAL		

Further analysis of the NOV38a protein yielded the following properties shown in Table 38B.

Table 38B. Protein Sequence Properties NOV38a	
PSort analysis:	0.6850 probability located in endoplasmic reticulum (membrane); 0.6400 probability located in plasma membrane; 0.4600 probability located in Golgi body; 0.1000 probability located in endoplasmic reticulum (lumen)
SignalP analysis:	Cleavage site between residues 27 and 28

A search of the NOV38a protein against the Geneseq database, a proprietary database that contains sequences published in patents and patent publication, yielded several homologous proteins shown in Table 38C.

Table 38C. Geneseq Results for NOV38a				
Geneseq Identifier	Protein/Organism/Length [Patent #, Date]	NOV38a Residues/ Match Residues	Identities/ Similarities for the Matched Region	Expect Value

AAR41876	Human HT6 - Homo sapiens, 230 aa. [DE4209216-A, 23-SEP-1993]	29..256 7..227	82/238 (34%) 120/238 (49%)	1e-21
AAW76806	Human ADP-ribosyltransferase protein - Homo sapiens, 327 aa. [US5834310-A, 10-NOV-1998]	20..266 31..287	83/266 (31%) 123/266 (46%)	6e-21
AAW76804	Rabbit skeletal muscle ADP-ribosyltransferase protein - Oryctolagus cuniculus, 327 aa. [US5834310-A, 10-NOV-1998]	8..259 6..280	88/282 (31%) 130/282 (45%)	1e-20
AAR37572	Rabbit skeletal muscle ADP-ribosyltransferase - Oryctolagus cuniculus, 327 aa. [USN7985698-N, 01-MAY-1993]	8..259 6..280	88/282 (31%) 130/282 (45%)	1e-20
ABB97573	Novel human protein SEQ ID NO: 841 - Homo sapiens, 229 aa. [WO200222660-A2, 21-MAR-2002]	29..163 29..161	59/137 (43%) 76/137 (55%)	1e-20

In a BLAST search of public sequence datbases, the NOV38a protein was found to have homology to the proteins shown in the BLASTP data in Table 38D.

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Table 38D. Public BLASTP Results for NOV38a				
Protein Accession Number	Protein/Organism/Length	NOV38a Residues/ Match Residues	Identities/ Similarities for the Matched Portion	Expect Value
S62906	mono-ADP-ribosyltransferase - human, 367 aa.	1..356 1..367	356/367 (97%) 356/367 (97%)	0.0
Q8WVJ7	Hypothetical 42.7 kDa protein - Homo sapiens (Human), 378 aa.	1..356 1..378	355/378 (93%) 355/378 (93%)	0.0
Q13508	Ecto-ADP-ribosyltransferase 3 precursor (EC 2.4.2.31) (NAD(P)(+)-- arginine ADP-ribosyltransferase 3) (Mono(ADP-ribosyl)transferas e 3) - Homo sapiens (Human), 389 aa.	1..356 1..389	355/389 (91%) 355/389 (91%)	0.0

| Q96HL1 | Unknown (protein for MGC:14489) - Homo sapiens (Human), 389 aa. | 1..356 1..389 | 354/389 (91%) 354/389 (91%) | 0.0 |
| Q9GKV6 | Hypothetical 38.2 kDa protein - Macaca fascicularis (Crab eating macaque) (Cynomolgus monkey), 338 aa. | 31..356 1..338 | 300/338 (88%) 312/338 (91%) | e-174 |

PFam analysis predicts that the NOV38a protein contains the domains shown in the Table 38E.

Table 38E. Domain Analysis of NOV38a			
Pfam Domain	NOV38a Match Region	Identities/ Similarities for the Matched Region	Expect Value
ART	1..312	164/340 (48%) 312/340 (92%)	1.5e-200

Example 39.

The NOV39 clone was analyzed, and the nucleotide and encoded polypeptide sequences are shown in Table 39A.

Table 39A. NOV39 Sequence Analysis			
	SEQ ID NO: 159	8350 bp	
NOV39a, CG150799-01 DNA Sequence	CAGGGAAAAGGGAACCTATGGAATGGTCATGGTGACTTTTGAGGTAGAGGGTGGCCCAAATCCCCCT GATGAAGATTTGAGTCCAGTTAAAGGAAATATCACCTTTCCCCCTGGCAGAGCAACAGTAATTTATA ACTTGACAGTACTCGATGACGAGGTACCAGAAAATGATGAAATATTTTTAATTCAACTGAAAAGTGT AGAAGGAGGAGCTGAGATTAACACCTCTAGGAATTCCATTGAGATCATCATTAAGAAAAATGATAGT CCCGTGAGATTCCTTCAGAGTATTTATTTGGTTCCTGAGGAAGACCACATACTCATAATTCCAGTAG TTCGTGGAAAGGACAACAATGGAAATCTGATTGGATCTGATGAATATGAGGTTTCAATCAGTTATGC TGTCACAACTGGGAATTCCACAGCACATGCCCAGCAAAATCTGGACTTCATTGATCTTCAGCCAAAC ACAACTGTTGTTTTTCCACCTTTTATTCATGAATCTCACTTGAAATTTCAAATAGTTGATGACACCA CACCGGAGATTGCTGAATCGTTTCACATTATGTTACTAAAAGATACCTTACAGGGAGATGCTGTGCT AATAAGCCCTTCTGTTGTACAAGTCACCATTAAGCCAAATGATAAACCTTATGGAGTCCTTTCATTC AACAGTGTTTTGTTTGAAAGGACAGTTATAATTGATGAAGATAGAATATCAAGATATGAAGAAATCA CAGTGGTTAGAAATGGAGGAACCCATGGGAATGTCTCTGCGAATTGGGTGTTGACACGGAACAGCAC TGATCCCTCACCAGTAACAGCAGATATCAGACCGAGCTCTGGAGTTCTCCATTTTGCACAAGGGCAG ATGTTGGCAACAATTCCTCTTACTGTGGTTGATGATGATCTTCCAGAAGAGGCAGAAGCTTATCTAC TTCAAATTCTGCCTCATACAATACGAGGAGGTGCAGAAGTGAGCGAGCCAGCGGAGGATAGTGATGA TGTCTATGGCCTAATAACATTTTTTCCTATGGAAAACCAGAAGATTGAAAGCAGCCCAGGTGAACGA TACTTATCCTTGAGTTTTACAAGACTAGGAGGGACTAAAGGAGATGTGAGGTTGCTTTATTCTGTAC TTTACATTCCTGCTGGAGCTGTGGACCCCTTGCAAGCAAAAGAAGGCATCTTAAATATATCAAGGAG AAATGACCTCATTTTTCCAGAGCAAAAAAACTCAAGTCACTACAAAATTACCAATAAGAAATGATGCA TTCTTTCAAAATGGAGCTCACTTTCTAGTACAGTTGGAAACTGTGGAGTTGTTAAACATAATTCCTC TAATCCCACCCATAAGCCCTAGATTTGGGGAAATCTGCAATATTTCTTTACTGGTTACTCCAGCCAT TGCAAATGGAGAAATTGGCTTTCTCAGCAATCTTCCAATTATTTTGCATGAACCAGAAGATTTTGCT GCTGAAGTGGTATACATTCCCTTACATCGGGATGGAACTGATGGCCAGGCTACTGTCTACTGGAGTT TGAAGCCCTCTGGCTTTAATTCAAAAGCAGTGACCCCGGATGATATAGGCCCCTTTAATGGCTCTGT

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AACAAACGAATCCCATGTGACACTTTCCCTTCATTATAAAACCTTGGGTTCCAATGCTACATACATT
GCCAAGACAACAGTCATGAAATATTTAGAAGAAAGTGTTTGGCTTCATCTACTAATTATCCTGGAGG
ATGGTATAATCGAATTCTACCTGGATGGAAATGACGCCAGGGAATCAAGAGTCTGAAAGGAGA
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ACTTAATGAGTATTTCCGTGTGACATTGGTTTCTGCAATTCCTGGAGATGGGAAGCTAGGCTCAACT
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ORF Start: ATG at 23		ORF Stop: TAA at 11537

	SEQ ID NO: 162	3838 aa	MW at 421384.3kD
NOV39b, CG150799-02 Protein Sequence	MVMVTFEVEGGPNPPDEDLSPVKGNITFPPGRATVIYNLTVLDDEVPENDEIFLIQLKSVEGGAEIN TSRNSIEIIIKKNDSPVRFLQSIYLVPEEDHILIIPVVRGKDNNGNLIGSDEYEVSISYAVTTGNST AHAQQNLDFIDLQPNTTVVFPPFIHESHLKFQIVDDTTPEIAESFHIMLLKDTLQGDAVLISPSVVQ VTIKPNDKPYGVLSFNSVLFERTVIIDEDRISRYEEITVVRNGGTHGNVSANWVLTRNSTDPSPVTA DIRPSSGVLHFAQGQMLATIPLTVVDDDLPEEAEAYLLQILPHTIRGGAEVSEPAEDSDDVYGLITF FPMENQKIESSPGERYLSLSFTRLGGTKGDVRLLYSVLYIPAGAVDPLQAKEGILNISRRNDLIFPE QKTQVTTKLPIRNDAFFQNGAHFLVQLETVELLNIIPLIPPISPRFGEICNISLLVTPAIANGEIGF LSNLPIIILHEPEDFAAEVVYIPLHRDGTDGQATVYWSLKPSGFNSKAVTPDDIGPFNGSVLFLSGQS DTTINITIKGDDIPEMNETVTLSLDRVNVENQVLKSGYTSRDLIILENDDPGGVFEFSPASRGPYVI KEGESVELHIIRSRGSLVKQFLHYRVEPRDSNEFYGNTGVLEFKPGEREIVITLLARLDGIPELDEH YWVVLSSHGERESKLGSATIVNITILKNDDPHGIIEFVSDGLIVMINESKGDAIYSAVYDVVRNRGN FGDVSVSWVVSPDFTQDVFPVQGTVVFGDQEFSKNITIYSLPDEIPEEMEEFTVILLNGTGGAKVGN RTTATLRIRRNDDPIYFAEPRVVRVQEGETANFTVLRNGSVDVTCMVQYATKDGKATARERDFIPVE KGETLIFEVGSRQQSISIFVNEDGIPETDEPFYIILLNSTGDTVVYQYGVATVIIEANDDPNGIFSL EPIDKAVEEGKTNAFWILRHRGYFGSVSVSWQLFQNDSALQPGQEFYETSGTVNFMDGEEAKPIILH AFPDKIPEFNEFYFLKLVNISGPGGQLAETNLQVTVMVPFNDDPFGVFILDPECLEREVAEDVLSED DMSYITNFTILRQQGVFGDVQLGWEILSSEFPAGLPPMIDFLLVGIFPTTVHLQQHMRRHHSGTDAL YFTGLEGAFGTVNPKYHPSRNNTIIANFTFSAWVMPNANTNGFIIAKDDGNGSIYYGVKIQTNESHVT LSLHYKTLGSNATYIAKTTVMKYLEESVWLHLLIILEDGIIEFYLDGNAMPRGIKSLKGEAITDGPG ILRIGAGINGNDRFTGLMQDVRSYERKLTLEEIYELHAMPAKSDLHPISGYLEFRQGETNKSFIISA RDDNDEEGEELFILKLVSVYGGARISEENTTARLTIQKSDNANGLFGFTGACIPEIAEEGSTISCVV ERTRGALDYVHVFYTISQIETDGINYLVDDFANASGTITFLPWQRSEVLNIYVLDDDIPELNEYFRV TLVSAIPGDGKLGSTPTSGASIDPEKETTDITIKASDHPYGLLQFSTGLPPQPKDAMTLPASSVPHI TVEEEDGEIRLLVIRAQGLLGRVTAEFRTVSLTAFSPEDYQNVAGTLEFQPGERYKYIFINITDNSI PELEKSFKVELLNLEGGVAELFRVDGSGSASLGVASQILVTIAASDHAHGVFEFSPESLFVSGTEPE DGYSTVTLNVIRHHGTLSPVTLHWNIDSDPDGLAFTSGNITFEIGQTSANITVEILPDEDPELDKA FSVSVLSVSSGSLGAHINATLTVLASDDPYGIFIFSEKNRPVKVEEATQNITLSIIRLKGLMGKVLV SYATLDDMEKPPYFPPNLARATOGRDYIPASGFALFGANOSEATIAISILDDDEPERSESVFIELLN

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	SEQ ID NO: 163	5102 bp	
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245

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	ORF Start: ATG at 23		ORF Stop: TGA at 4430

	SEQ ID NO: 164	1469 aa	MW at 162809.6kD
NOV39c, CG150799-03 Protein Sequence	MVMVTFEVEGGGPNPPDEDLSPVKGNITFPPGRATVIYNLTVLDDEVPENDEIFLIQLKSVEGGAEIN TSRNSIEIIIKKNDSPVRFLQSIYLVPEEDHILIIPVVRGKDNNGNLIGSDEYEVSISYAVTTGNST AHAQQNLDFIDLQPNTTVVFPPFIHESHLKFQIVDDTTPEIAESFHIMLLKDTLQGDAVLISPSVVQ VTIKPNDKPYGVLSFNSVLFERTVIIDEDRISRYEEITVVRNGGTHGNVSANWVLTRNSTDPSPVTA DIRPSSGVLHFAQGQMLATIPLTVVDDDLPEEAEAYLLQILPHTIRGGAEVSEPAEDSDDVYGLITF FPMENQKIESSPGERYLSLSFTRLGGTKGDVRLLYSVLYIPAGAVDPLQAKEGILNISRRNDLIFPE QKTQVTTKLPIRNDAFFQNGAHFLVQLETVELLNIIPLIPPISPRFGEICNISLLVTPAIANGEIGF LSNLPIIILHEPEDFAAEVVYIPLHRDGTDGQATVYWSLKPSGFNSKAVTPDDIGPFNGSVLFLSGQS DTTINITIKGDDIPEMNETVTLSLDRVNVENQVLKSGYTSRDLIILENDDPGGVFEFSPASRGPYVI KEGESVELHIIRSRGSLVKQFLHYRVEPRDSNEFYGNTGVLEFKPGEREIVITLLARLDGIPELDEH YWVVLSSHGERESKLGSATIVNITILKNDDPHGIIEFVSDGLIVMINESKGDAIYSAVYDVVRNRGN FGDVSVSWVVSPDFTQDVFPVQGTVVFGDQEFSKNITIYSLPDEIPEEMEEFTVILLNGTGGAKVGN RTTATLRIRRNDDPIYFAEPRVVRVQEGETANFTVLRNGSVDVTCMVQYATKDGKATARERDFIPVE KGETLIFEVGSRQQSISIFVNEDGIPETDEPPFYIILLNSTGDTVVYQYGVATVIIEANDDPNGIFSL EPIDKAVEEGKTNAFWILRHRGYFGSVSVSWQLFQNDSALQPGQEFYETSGTVNFMDGEEAKPIILH AFPDKIPEFNEFYFLKLVNISGPGGQLAETNLQVTVMVPFNDDPFGVFILDPECLEREVAEDVLSED DMSYITNFTILRQQGVFGDVQLGWEILSSEFPAGLPPMIDFLLVGIFPTTVHLQQHMRRHHSGTDAL YFTGLEGAFGTVNPKYHPSRNNTIANFTFSAWVMPNANTNGFIIAKDDGNGSIYYGVKIQTNESHVT LSLHYKTLGSNATYIAKTTVMKYLEESVWLHLLIILEDGIIEFYLDGNAMPRGIKSLKGEAITDGPG ILRIGAGINGNDRFTGLMQDVRSYERKLTLEEIYELHAMPAKSDLHPISGYLEFRQGETNKSPIISA RDDNDEEGEELFILKLVSVYGGARISEENTTARLTIQKSDNANGLFGFTGACIPEIAEEGSTISCVV ERTRGALDYVHVFYTISQIETDGINYLVDDFANASGTITFLPWQRSELLIEVSLPIIIYNCN

	SEQ ID NO: 165	8350 bp	
NOV39d, CG150799-01 DNA Sequence	<u>CAGGGAAAAGGGAACC</u>TATGG<u>A</u>ATGGTCATGGTGACTTTTGAGGTAGAGGGTGGCCCAAATCCCCCT GATGAAGATTTGAGTCCAGTTAAAGGAAATATCACCTTTCCCCCTGGCAGAGCAACAGTAATTTATA ACTTGACAGTACTCGATGACGAGGTACCAGAAATGATGAAATATTTTTAATTCAACTGAAAAGTGT AGAAGGAGGAGCTGAGATTAACACCTCTAGGAATTCCATTGAGATCATCATTAAGAAAAATGATAGT CCCGTGAGATTCCTTCAGAGTATTTATTTGGTTCCTGAGGAAGACCACATACTCATAATTCCAGTAG TTCGTGGAAAGGACAACAATGGAAATCTGATTGGATCTGATGAATATGAGGTTTCAATCAGTTATGC TGTCACAACTGGGAATTCCACAGCACATGCCCAGCAAAATCTGGACTTCATTGATCTTCAGCCAAAC ACAACTGTTGTTTTTCCACCTTTTATTCATGAATCTCACTTGAAATTTCAAATAGTTGATGACACCA CACCGGAGATTGCTGAATCGTTTCACATTATGTTACTAAAAGATACCTTACAGGGAGATGCTGTGCT AATAAGCCCTTCTGTTGTACAAGTCACCATTAAGCCAAATGATAAACCTTATGGAGTCCTTTCATTC AACAGTGTTTTGTTTGAAAGGACAGTTATAATTGATGAAGATAGAATATCAAGATATGAAGAAATCA CAGTGGTTAGAAATGGAGGAACCCATGGGAATGTCTCTGCGAATTGGGTGTTGACACGGAACAGCAC TGATCCCTCACCAGTAACAGCAGATATCAGACCGAGCTCTGGAGTTCTCCATTTTGCACAAGGGCAG ATGTTGGCAACAATTCCTCTTACTGTGGTTGATGATGATCTTCCAGAAGAGGCAGAAGCTTATCTAC TTCAAATTCTGCCTCATCACAATACGAGGAGGTGCAGAAGTGAGCGAGCCAGCGGAGGATAGTGATGA TGTCTATGGCCTAATAACATTTTTTCCTATGGAAAACCAGAAGATTGAAAGCAGCCCAGGTGAACGA TACTTATCCTTGAGTTTTACAAGACTAGGAGGGACTAAAGGAGATGTGAGGTTGCTTTATTCTGTAC TTTACATTCCTGCTGGAGCTGTGGACCCCTTGCAAGCAAAAGAAGGCATCTTAAATATATCAAGGAG AAATGACCTCATTTTTCCAGAGCAAAAATTACCAAGTCACTACAAAATTACCAATAAGAAATGATGCA TTCTTTCAAAATGGAGCTCACTTTCTAGTACAGTTGGAAACTGTGGAGTTGTTAAACATAATTCCTC TAATCCCACCCATAAGCCCTAGATTTGGGGAAATCTGCAATATTTCTTTACTGGTTACTCCAGCCAT TGCAAATGGAGAAATTGGCTTTCTCAGCAATCTTCCAATTATTTTGCATGAACCAGAAGATTTTGCT GCTGAAGTGGTATACATTCCCTTACATCGGGATGGAACTGATGGCCAGGCTACTGTCTACTGGAGTT TGAAGCCCTCTGGCTTTAATTCAAAAGCAGTGACCCCGGATGATATATAGGCCCCTTTAATGGCTCTGT TTTGTTTTTATCTGGGCAAAGTGACACAACAATCAACATTACTATCAAAGGTGATGACATACCGGAA ATGAATGAAACTGTAACACTTTCTCTAGACAGGGTTAACGTGGAAAACCAAGTGCTGAAATCTGGAT ATACTAGCCGTGACCTAATTATTTTGGAAAATGATGACCCTGGGGGAGTTTTTGAATTTTCTCCTGC TTCCAGAGGACCCTATGTTATAAAAGAAGGAGAATCTGTAGAGCTCCACATCATCCGATCAAGGGGG TCCCTTGTTAAGCAGTTTCTACACTACCGAGTAGAGCCAAGAGATAGCAATGAATTCTATGGAAACA CGGGAGTACTAGAATTTAAACCTGGAGAAAGGGAGATAGTGATCACCTTGCTAGCAAGATTGGATGG GATACCAGAGTTGGATGAACACTACTGGGTGGTCCTCAGCAGCCACGGAGAACGGGAAAGCAAGTTG GGAAGTGCCACCATTGTCAATATAACGATTCTGAAAAATGATGATCCTCATGGCATTATAGAATTTG TTTCTGATGGTCTAATTGTGATGATAAATGAAAGCAAAGGAGATGCTATCTATAGTGCTGTTTATGA TGTAGTAAGAAATCGAGGCAACTTTGGTGATGTTAGTGTATCATGGGTGGTTAGTCCAGACTTTACA CAAGATGTATTTCCTGTACAAGGGACTGTTGTCTTTGGAGATCAGGAATTTTCAAAAAATATCACCA TTTACTCCCTTCCAGATGAGATTCCAGAAGAAATGGAAGAATTTACCGTTATCCTACTGAATGGCAC TGGAGGAGCTAAAGTGGGAAATAGAACAACTGCAACTCTGAGGATTAGAAGAAATGATGACCCCATT TATTTTGCAGAACCTCGTGTAGTGAGGGTTCAGGAAGGTGAGACTGCCAACTTTACAGTTCTCAGAA ATGGATCTGTTGATGTGACTTGCATGGTCCAGTATGCTACCAAGGATGGGAAGGCTACTGCAAGAGA GAGAGATTTCATTCCTGTTGAAAAAGGAGAAACGCTCATTTTTGAGGTTGGAAGTAGACAGCAGAGC ATATCCATATTTGTTAATGAAGATGGTATCCCGGAAACAGATGAGCCCTTTTATATAATCCTCTTGA ATTCAACAGGTGATACAGTAGTATATCAATATGGAGTAGCTACAGTAATAATTGAAGCTAATGATGA CCCAAATGGCATTTTTTTCTCTGGAGCCCATAGACAAAGCAGTGGAAGAAGGAAAGACTAATGCATTT TGGATTTTGAGGCACCGAGGATACTTTGGTAGTGTTTCTGTATCTTGGCAGCTCTTTCAGAATGATT CTGCTTTGCAGCCTGGGCAGGAGTTCTATGAAACTTCAGGAACTGTTAACTTCATGGATGGAGAAGA AGCAAAACCAATCATTCTCCATGCTTTTTCCAGATAAAATTCCTGAATTCAATGAATTTTATTTCCTA AAACTTGTAAACATTTCAGGTCCTGGGGGCCAGCTAGCAGAAACCAACCTCCAGGTGACAGTAATGG TTCCATTCAATGATGATCCCTTTGGAGTTTTTATCTTGGATCCAGAGTGTTTAGAGAGAGAAGTGGC AGAAGATGTCCTGTCTGAAGATGATATGTCTTATATTACCAACTTCACCATTTTGAGGCAGCAGGGT GTGTTTGGTGATGTACAACTGGGCTGGGAAATACTGTCCAGTGAGTTCCCTGCTGGTTTGCCACCAA TGATAGATTTTTTTACTGGTTGGAATTTTCCCCACCACCGTGCATTTACAACAGCACATGCGGCGTCA CCACAGTGGAACGGATGCTTTGTACTTTACCGGACTAGAGGGTGCATTTGGGACTGTTAATCCAAAA TACCATCCCTCCAGGAATAATACAATTGCCAACTTTACATTCTCAGCTTGGGTAATGCCCAATGCCA ATACGAATGGATTCATTATAGCGAAGGATGACGGTAATGGAAGCATCTACTACGGGTAAAAATACA AACAAACGAATCCCATGTGACACTTTCCCTTCATTATAAAACCTTGGGTTCCAATGCTACATACATT GCCAAGACAACAGTCATGAAATATTTAGAAGAAAGTGTTTGGCTTCATCTACTAATTATCCTGGAGG ATGGTATAATCGAATTCTACCTGGATGGAAATGCAATGCCCAGGGGAATCAAGAGTCTGAAAGGAGA AGCCATTACTGACGGTCCTGGGATACTGAGAATTGGAGCAGGGATAAATGGCAATGACAGATTTACA GGTCTGATGCAGGATGTGAGGTCCTATGAGCGGAAACTGACGCTTGAAGAAATTTATGAACTTCATG CCATGCCCGCAAAAAGTGATTTACACCCAATTTCTGGATATCTGGAGTTCAGACAGGGAGAAACTAA CAAATCATTCATTATTTCTGCAAGACGATGACAATGACGAGGAAGGAGAAGAATTATTCATTCTTAAA CTAGTTTCTGTATATGGAGGAGCTCGTATTTCGGAAGAAAATACTACTGCAAGATTAACAATACAAA AAAGTGACAATGCAAATGGCTTGTTTGGTTTCACAGGAGCTTGTATACCAGAGATTGCAGAGGAGGG ATCAACCATTTCTTGTGTGGTTGAGAGAACCAGAGGAGCTCTGGATTATGTGCATGTTTTTTACACC ATTTCACAGATTGAAACTGATGGCATTAATTACCTTGTTGATGACTTTGCTAATGCCAGTGGAACTA TTACATTCCTTCCTTGGCAGAGATCAGAGGTTCTGAATATATATGTTCTTGATGATGATATTCCTGA ACTTAATGAGTATTTCCGTGTGACATTGGTTTCTGCAATTCCTGGAGATGGGAAGCTAGGCTCAACT CCTACCAGTGGTGCAAGCATAGATCCTGAAAAGGAAACGACTGATATCACCATCAAAGCTAGTGATC ATCCATATGGCTTGCTGCAGTTCTCCACAGGGCTGCCTCCTCAGCCTAAGGACGCAATGACCCTGCC TGCAAGCAGCCGTTCCACATATCACTGTGGAGGAGAATGGAGAAATCAGGTTATTGGTCATCCGT GCACAGGGACTTCTGGGAAGGGTGACTGCGGAATTTAGAACAGTGTCCTTGACAGCATTCAGTCCTG AGGATTACCAGAATGTTGCTGGCACATTAGAATTTCAACCAGGAGAAAGATATAAATACATTTTTCAT

| AAACATCACTGATAATTCTATTCCTGAACTGGAAAATCTTTTAAAGTTGAGTTGTTAAACTTGGAA |
| GGAGGAGTAGCTGAACTCTTTAGGGTTGATGGAAGTGGTAGTGCCAGTCTAGGAGTGGCTTCCCAAA |
| TTCTAGTGACAATTGCAGCCTCTGACCAGCTCATGGCGTATTTGAATTTAGCCCTGAGTCACTCTT |
| TGTCAGTGGAACTGAACCAGAAGATGGGTATAGCACTGTTACATTAAATGTTATAAGACATCATGGA |
| ACTCTGTCTCCAGTGACTTTGCATTGGAACATAGACTCTGATCCTGATGGTGATCTCGCCTTCACCT |
| CTGGCAACATCACATTTGAGATTGGGCAGACGAGCGCCAATATCACTGTGGAGATATTGCCTGACGA |
| AGACCCGAACTGGATAAGGCATTCTCTGTGTCAGTCCTCAGTGTTTCCAGTGGTTCTTTGGGAGCT |
| CATATTAATGCCACGTTAACAGTTTTGGCTAGTGATGATCCATATGGGATATTCATTTTTTCTGAGA |
| AAAACAGACCTGTTAAAGTTGAGGAAGCAACCCAGAACATCACACTATCAATAATAAGGTTGAAAGG |
| CCTCATGGGAAAAGTCCTTGTCTCATATGCAACACTAGATGATATGGAAAAACCACCTTATTTTCCA |
| CCTAATTTAGCGAGAGCAACTCAAGGAAGAGACTATATACCAGCTTCTGGATTTGCTCTTTTTGGAG |
| CTAATCAGAGTGAGGCAACAATAGCTATTTCAATTTTGGATGATGATGAGCCAGAAAGGTCCGAATC |
| TGTCTTTATCGAACTACTCAACTCTACTTTAGTAGCGAAAGTACAGAGTCGTTCAATTCCAAATTCT |
| CCACGTCTTGGGCCTAAGGTAGAAACTATTGCGCAACTAATTATCATTGCCAATGATGATGCATTTG |
| GAACTCTTCAGCTCTCAGCACCAATTGTCCGAGTGGCAGAAAATCATGTTGGACCCATTATCAATGT |
| GACTAGAACAGGAGGAGCATTTGCAGATGTCTCTGTGAAGTTTAAAGCTGTGCCAATAACTGCAATA |
| GCTGGTGAAGATTATAGTATAGCTTCATCAGATGTGGTCTTGCTAGAAGGGGAAACCAGTAAAGCCG |
| TGCCAATATATGTCATTAATGATATCTATCCTGAACTGGAAGAATCTTTTCTTGTGCAACTGATGAA |
| TGAAACAACAGGAGGAGCCAGACTAGGGGCTTTAACAGAGGCAGTCATTATTATTGAGGCCTCTGAT |
| GACCCCTATGGATTATTTGGTTTTCAGATTACTAAACTTATTGTAGAGGAACCTGAGTTTAACTCAG |
| TGAAGGTAAACCTGCCAATAATTCGAAATTCTGGGACACTCGGCAATGTTACTGTTCAGTGGGTTGC |
| CACCATTAATGGACAGCTTGCTACTGGCGACCTGCGAGTTGTCTCAGGTAATGTGACCTTTGCCCCT |
| GGGGAAACCATTCAAACCTTGTTGTTAGAGGTCCTGGCTGACGACGTTCCGGAGATTGAAGAGGTTA |
| TCCAAGTGCAACTAACTGATGCCTCTGGTGGAGGTACTATTGGGTTAGATCGAATTGCAAATATTAT |
| TATTCCTGCCAATGATGATCCTTATGGTACAGTAGCCTTTGCTCAGATGGTTTATCGTGTTCAAGAG |
| CCTCTGGAAGAAGTTCCTGTGCTAATATAACTGTCAGGCGAAGCGGAGGGCACTTTGGTCGGCTGT |
| TGTTGTTCTACAGTACTTCCGACATTGATGTAGTGGCTCTGGCAATGGAGGAAGGTCAAGATTTACT |
| GTCCTACTATGAATCTCCAATTCAAGGGGTGACCTCACCTTTGGAGAACTTGGATGAATGTCTCT |
| GCCGTGGGGGAGCCCCTGTATACCTGTGCCACTTTGTGCCTTAAGGAACAAGCTTGCTCAGCGTTTT |
| CATTTTTCAGTGCTTCTGAGGGTCCCCAGTGTTTCTGGATGACATCATGGATCAGCCCAGCTGTCAA |
| CAATTCAGACTTCTGGACCTACAGGAAAAACATGACCAGGGTAGCATCTCTTTTTAGTGGTCAGGCT |
| GTGGCTGGGAGTGACTATGAGCCTGTGACAAGGCAATGGGCCATAATGCAGGAAGGTGATGAATTCG |
| CAAATCTCACAGTGTCTATTCTTCCTGATGATTTCCCAGAGATGGATGAGAGTTTTCTAATTTCTCT |
| CCTTGAAGTTCACCTCATGAACATTTCAGCCAGTTTGAAAAATCAGCCAACCATAGGACAGCCAAAT |
| ATTTCTACAGTTGTCATAGCACTAAATGGTGATGCCTTTGGAGTGTTTGTGATCTACAATATTAGTC |
| CCAATACTTCCGAAGATGGCTTATTTGTTGAAGTTCAGGAGCAGCCCCAAACCTTGGTGGAGCTGAT |
| GATACACAGGACAGGGGGCAGCTTAGGTCAAGTGGCAGTCGAATGGCGTGTTGTTGGTGGAACAGCT |
| ACTGAAGGTTTAGATTTTATAGGTGCTGGAGAGATTCTGACCTTTGCTGAAGGTGAAACCAAAAAGA |
| CAGTCATTTTAACCATCTTGGATGACTCTGAACCAGAGGATGACGAAAGTATCATAGTTAGTTTGGT |
| GTACACTGAAGGTGGAAGTAGAATTTTGCCAAGCTCCGACACTGTTAGAGTGAACATTTTGGCCAAT |
| GACAATGTGGCAGGAATTGTTAGCTTTCAGACAGCCTTCCAGATCTGTCATAGGTCATGAAGGAGAAA |
| TTTTACAATTCCATGTGATAAGAACTTTCCCTGGTCGAGGAAATGTTACTGTTAACTGGAAAATTAT |
| TGGGCAAAATCTAGAACTCAATTTTGCTAACTTTAGCGGACAACTTTTCTTTCCTGAGGGGTCGTTG |
| AATACAACATTGTTTGTGCATTTGTTGGATGACAACATTCCTGAGGAGAAAGAAGTATACCAAGTCA |
| TTCTGTATGATGTCAGGACACAAGGAGTTCCACCAGCCGGAATCGCCCCTGCTTGATGCTCAAGGATA |
| TGCAGCTGTCCTCACAGTAGAAGCCAGTGATGAACCACATGGAGTTTTAAATTTTGCTCTTTCATCA |
| AGATTTGTGTTACTACAAGAGGCTAACATAACAATTCAGCTTTTCATCAACAGAGAATTTGGATCTC |
| TAGGAGCTATCAATGTCACATATACCACGGTTCCTGGAATGCTGAGTCTGAAGAACCAAACAGTAGG |
| AAACCTAGCAGAGCCAGAAGTTGATTTTGTCCCTATCATTGGCTTTCTGATTTTAGAAGAAGGGGAA |
| ACAGCAGCAGCCATCAACATTACCATTCTTGAGGATGATGTACCAGAGCTAGAAGAATATTTCCTGG |
| TGAATTTAACTTACGTTGGACTTACCATGGCTGCTTCAACTTCATTTCCTCCCAGACTAGGTATGAG |
| GGGTTTCTTGTTTGTTTCTTTTTGCTCACTTCAAATGAAATGAAGAAACTTCATTTTTGAATCAGAA |
| GTGATCATTGTGCTGTTTTGTTAATCTTAGCTATGTGTTAAA |

| ORF Start: ATG at 23 | | | ORF Stop: TGA at 8282 |

	SEQ ID NO: 166	2753 aa	MW at 301743.8kD
NOV39d, CG150799-01 Protein Sequence	MVMVTFEVEGGPNPPDEDLSPVKGNITFPPGRATVIYNLTVLDDEVPENDEIFLIQLKSVEGGAEIN TSRNSIEIIIKKNDSPVRFLQSIYLVPEEDHILIIPVVRGKDNNGNLIGSDEYEVSISYAVTTGNST AHAQQNLDFIDLQPNTTVVFPPPFIHESHLKFQIVDDTTPEIAESFHIMLLKDTLQGDAVLISPSVVQ VTIKPNDKPYGVLSFNSVLFERTVIIDEDRISRYEEITVVRNGGTHGNVSANWVLTRNSTDPSPVTA DIRPSSGVLHFAQGQMLATIPLTVVDDDLPEEAEAYLLQILPHTIRGGAEVSEPAEDSDDVYGLITF FPMENQKIESSPGERYLSLSFTRLGGTKGDVRLLYSVLYIPAGAVDPLQAKEGILNISRRNDLIFPE QKTQVTTKLPIRNDAFFQNGAHFLVQLETVELLNIIPLIPPISPRFGEICNISLLVTPAIANGEIGF LSNLPIIILHEPEDFAAEVVYIPLHRDGTDGQATVYWSLKPSGFNSKAVTPDDIGPFNGSVLFLSGQS DTTINITIKGDDIPEMNETVTLSLDRVNVENQVLKSGYTSRDLIILENDDPGGVFEFSPASRGPYVI KEGESVELHIIRSRGSLVKQFLHYRVEPRDSNEFYGNTGVLEFKPGEREIVITLLARLDGIPELDEH YWVVLSSHGERESKLGSATIVNITILKNDDPHGIIEFVSDGLIVMINESKGDAIYSAVYDVVRNRGN FGDVSVSWVVSPDFTODVFPVOGTVVFGDOEFSKNITIYSLPDEIPEEMEEFTVILLNGTGGAKVGN

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RTTATLRIRRNDDPIYFAEPRVVRVQEGETANFTVLRNGSVDVTCMVQYATKDGKATARERDFIPVE
KGETLIPEVGSRQQSISIFVNEDGIPETDEPFYIILLNSTGDTVVYQYGVATVIIEANDDPNGIFSL
EPIDKAVEEGKTNAFWILRHRGYFGSVSVSWQLFQNDSALQPGQEFYETSGTVNFMDGEEAKPIILH
AFPDKIPEFNEFYFLKLVNISGPGGQLAETNLQVTVMVPFNDDPFGVFILDPECLEREVAEDVLSED
DMSYITNFTILRQQGVFGDVQLGWEILSSEFPAGLPPMIDFLLVGIFPTTVHLQQHMRRHHSGTDAL
YFTGLEGAFGTVNPKYHPSRNNTIANFTFSAWVMPNANTNGFIIAKDDGNGSIYYGVKIQTNESHVT
LSLHYKTLGSNATYIAKTTVMKYLEESVWLHLLIILEDGIIEFYLDGNAMPRGIKSLKGEAITDGPG
ILRIGAGINGNDRFTGLMQDVRSYERKLTLEEIYELHAMPAKSDLHPISGYLEFRQGETNKSFIISA
RDDNDEEGEELFILKLVSVYGGARISEENTTARLTIQKSDNANGLFGFTGACIPEIAEEGSTISCVV
ERTRGALDYVHVFYTISQIETDGINYLVDDFANASGTITFLPWQRSEVLNIYVLDDDIPELNEYFRV
TLVSAIPGDGKLGSTPTSGASIDPEKETTDITIKASDHPYGLLQFSTGLPPQPKDAMTLPASSVPHI
TVEEEDGEIRLLVIRAQGLLGRVTAEFRTVSLTAFSPEDYQNVAGTLEFQPGERYKYIFINITDNSI
PELEKSFKVELLNLEGGVAELFRVDGSGSASLGVASQILVTIAASDHAHGVFEFSPESLFVSGTEPE
DGYSTVTLNVIRHHGTLSPVTLHWNIDSDPDGDLAFTSGNITFEIGQTSANITVEILPDEDPELDKA
FSVSVLSVSSGSLGAHINATLTVLASDDPYGIFIFSEKNRPVKVEEATQNITLSIIRLKGLMGKVLV
SYATLDDMEKPPYFPPNLARATQGRDYIPASGFALFGANQSEATIAISILDDDEPERSESVFIELLN
STLVAKVQSRSIPNSPRLGPKVETIAQLIIIANDDAFGTLQLSAPIVRVAENHVGPIINVTRTGGAF
ADVSVKFKAVPITAIAGEDYSIASSDVVLLEGETSKAVPIYVINDIYPELEESFLVQLMNETTGGAR
LGALTEAVIIIEASDDPYGLFGFQITKLIVEEPEFNSVKVNLPIIRNSGTLGNVTVQWVATINGQLA
TGDLRVVSGNVTFAPGETIQTLLLEVLADDVPEIEEVIQVQLTDASGGGTIGLDRIANIIIPANDDP
YGTVAFAQMVYRVQEPLERSSCANITVRRSGGHFGRLLLFYSTSDIDVVALAMEEGQDLLSYYESPI
QGVPDPLWRTWMNVSAVGEPLYTCATLCLKEQACSAFSFFSASEGPQCFWMTSWISPAVNNSDFWTY
RKNMTRVASLFSGQAVAGSDYEPVTRQWAIMQEGDEFANLTVSILPDDPPEMDESFLISLLEVHLMN
ISASLKNQPTIGQPNISTVVIALNGDAFGVFVIYNISPNTSEDGLFVEVQEQPQTLVELMIHRTGGS
LGQVAVEWRVVGGTATEGLDFIGAGEILTFAEGETKKTVILTILDDSEPEDDESIIVSLVYTEGGSR
ILPSSDTVRVNILANDNVAGIVSFQTASRSVIGHEGEILQFHVIRTFPGRGNVTVNWKIIGQNLELN
FANFSGQLFFPEGSLNTTLFVHLLDDNIPEEKEVYQVILYDVRTQGVPPAGIALLDAQGYAAVLTVE
ASDEPHGVLNFALSSRFVLLQEANITIQLFINREFGSLGAINVTYTTVPGMLSLKNQTVGNLAEPEV
DFVPIIGFLILEEGETAAAINITILEDDVPELEEYFLVNLTYVGLTMAASTSFPPRLGMRGFLFVSF
CSLQMK
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Sequence comparison of the above protein sequences yields the following sequence relationships shown in Table 39B.

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Table 39B. Comparison of NOV39a against NOV39b through NOV39d.		
Protein Sequence	NOV39a Residues/ Match Residues	Identities/ Similarities for the Matched Region
NOV39b	1..2741 1..2741	2684/2741 (97%) 2685/2741 (97%)
NOV39c	1..1456 1..1456	1442/1456 (99%) 1443/1456 (99%)
NOV39d	1..2753 1..2753	2700/2753 (98%) 2700/2753 (98%)

Further analysis of the NOV39a protein yielded the following properties shown in Table 39C.

10 Table 39C.

Table 39C. Protein Sequence Properties NOV39a

PSort analysis:	0.5050 probability located in cytoplasm; 0.3836 probability located in microbody (peroxisome); 0.1851 probability located in lysosome (lumen); 0.1000 probability located in mitochondrial matrix space
SignalP analysis:	No Known Signal Sequence Predicted

A search of the NOV39a protein against the Geneseq database, a proprietary database that contains sequences published in patents and patent publication, yielded several homologous proteins shown in Table 39D.

Table 39D. Geneseq Results for NOV39a				
Geneseq Identifier	Protein/Organism/Length [Patent #, Date]	NOV39a Residues/ Match Residues	Identities/ Similarities for the Matched Region	Expect Value
AAE10925	Human monogenic audiogenic seizure-susceptible-1 (mass1) protein - Homo sapiens, 2777 aa. [WO200165927-A1, 13-SEP-2001]	1..2753 1..2777	2736/2778 (98%) 2739/2778 (98%)	0.0
AAE10924	Mouse monogenic audiogenic seizure-susceptible-1 (mass1) protein - Mus musculus, 2780 aa. [WO200165927-A1, 13-SEP-2001]	1..2739 1..2761	2295/2762 (83%) 2516/2762 (91%)	0.0
AAE10949	Mouse mass1 protein mutant (7009deltaG) - Mus musculus, 2071 aa. [WO200165927-A1, 13-SEP-2001]	1..2049 1..2071	1710/2072 (82%) 1878/2072 (90%)	0.0
ABG61545	Human transporter and ion channel, TRICH15, Incyte ID 7476089CD1 - Homo sapiens, 759 aa. [WO200240541-A2, 23-MAY-2002]	1531..2288 1..746	740/758 (97%) 740/758 (97%)	0.0
ABB05663	Human signal transduction protein clone amy2_10p7 - Homo sapiens, 1615 aa. [WO200198454-A2, 27-DEC-2001]	2232..2741 9..518	506/510 (99%) 507/510 (99%)	0.0

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In a BLAST search of public sequence datbases, the NOV39a protein was found to have homology to the proteins shown in the BLASTP data in Table 39E.

5

Table 39E. Public BLASTP Results for NOV39a				
Protein Accession Number	Protein/Organism/Length	NOV39a Residues/ Match Residues	Identities/ Similarities for the Matched Portion	Expect Value
Q8WXG9	Very large G protein-coupled receptor 1b - Homo sapiens (Human), 6307 aa.	1..2741 180..2945	2721/2766 (98%) 2723/2766 (98%)	0.0
Q91ZS2	MASS1 - Mus musculus (Mouse), 2780 aa.	1..2739 1..2761	2293/2762 (83%) 2515/2762 (91%)	0.0
Q8VHN7	Very large G protein-coupled receptor 1 - Mus musculus (Mouse), 6298 aa.	1..2741 179..2941	2293/2764 (82%) 2514/2764 (89%)	0.0
Q91ZS1	MASS1.2 - Mus musculus (Mouse), 2238 aa.	563..2739 29..2219	1838/2192 (83%) 2004/2192 (90%)	0.0
Q8TF58	KIAA1943 protein - Homo sapiens (Human), 1054 aa (fragment).	234..1273 1..1050	1037/1050 (98%) 1037/1050 (98%)	0.0

Example 40.

10   The NOV40 clone was analyzed, and the nucleotide and encoded polypeptide sequences are shown in Table 40A.

Table 40A. NOV40 Sequence Analysis			
	SEQ ID NO: 167	2833 bp	
NOV40a, CG151014-01 DNA Sequence	CAAAGATCCAGTTTGGAAATGAGAGAGGACTAGCATGACACATTGGCTCCACCATTGATATCTCCCA GAGGTACAGAAACAGGATTCATGAAGATGTTGACAAGACTGCAAGTTCTTACCTTAGCTTTGTTTTC AAAGGGATTTTTACTCTCTTTAGGGGACCATAACTTTCTAAGGAGAGAGATTAAAATAGAAGGTGAC CTTGTTTTAGGGGGCCTGTTTCCTATTAACGAAAAAGGCACTGGAACTGAAGAATGTGGGCGAATCA ATGAAGACCGAGGGATTCAACGCCTGGAAGCCATGTTGTTTGCTATTGATGAAATCAACAAAGATGA TTACTTGCTACCAGGAGTGAAGTTGGGTGTTCACATTTTGGATACATGTTCAAGGGATACCTATGCA TTGGAGCAATCACTGGAGTTTGTCAGGGCATCTTTGACAAAAGTGGATGAAGCTGAGTATATGTGTC CTGATGGATCCTATGCCATTCAAGAAAACATCCCACTTCTCATTGCAGGGGTCATTGGTGGCTCTTA TAGCAGTGTTTCCATACAGGTGGCAAACCTGCTGCGGCTCTTCCAGATCCCTCAGATCAGCTACGCA TCCACCAGCGCCAAACTCAGTGATAAGTCGCGCTATGATTACTTTGCCAGGACCGTGCCCCCCGACT TCTACCAGGCCAAAGCCATGGCTGAGATCTTGCGCTTCTTCAACTGGACCTACGTGTCCACAGTAGC CTCCGAGGGTGATTACGGGGGAGACAGGGATCGAGGCCTTCGAGCAGGAAGCCCGCCTGCGCAACATC TGCATCGCTACGGCGGAGAAGGTGGGCCGCTCCAACATCCGCAAGTCCTACGACAGCGTGATCCGAG AACTGTTGCAGAAGCCCAACGCGCGCGTCGTGGTCCTCTTCATGCGCAGCGACGACTCGCGGGAGCT CATTGCAGCCGCCAGCCGCGCCAATGCCTCCTTCACCTGGGTGGCCAGCGACGGCTGGGGCGCGCAG GAGAGCATCATCAAGGGCAGCGAGCATGTGGCCTACGGCGCCATCACCCTGGAGCTGGCCTCCCAGC

CTGTCCGCCAGTTCGACCGCTACTTCCAGAGCCTCAACCGCTACAACAACCACGGCAACGGCTGGTTC CCGGGACTTCTGGGAGCAAAAGTTTCAGTGCAGCCTCCAGAACAAACGCAACCACAGGCGCGTCTGC GACAAGCACCTGGCCATCGACGAGCAGCAACTACGAGCAAGAGTCCAAGATCATGTTTGTGGTGAACG CGGTGTATGCCATGGCCCACGCTTTGCACAAAATGCAGCGCACCCTCTGTCCCAACACTACCAAGCT TTGTGATGCTATGAAGATCCTGGATGGGAAGAAGTTGTACAAGGATTACTTGCTGAAAATCAACTTC ACGGCTCCATTCAACCCAAATAAAGATGCAGATAGCATAGTCAAGTTTGACACTTTTGGAGATGGAA TGGGGCGATACAACGTGTTCAATTTCCAAAATGTAGGTGGAAAGTATTCCTACTTGAAAGTTGGTCA CTGGGCAGAAACCTTATCGCTAGATGTCAACTCTATCCACTGGTCCCGGAACTCAGTCCCCACTTCC CAGTGCAGCGACCCCTGTGCCCCCAATGAAATGAAGAATATGCAACCAGGGGATGTCTGCTGCTGGA TTTGCATCCCCTGTGAACCCTACGAATACCTGGCTGATGAGTTTACCTGTATGGATTGTGGGTCTGG ACAGTGGCCCACTGCAGACCTAACTGGATGCTATGACCTTCCTGAGGACTACATCAGGTGGGAAGAC GCCTGGGCCATTGGCCCAGTCACCATTGCCTGTCTGGGTTTTATGTGTACATGCATGGTTGTAACTG TTTTTATCAAGCACAACAACACACCCTTGGTCAAAGCATCGGGCCGAGAACTCTGCTACATCTTATT GTTTGGGGGTTGGCCTGTCATACTGCATGACATTCTTCTTCATTGCCAAGCCATCACCAGTCATCTGT GCATTGCGCCGACTCGGGCTGGGGAGTTCCTTCGCTATCTGTTACTCAGCCCTGCTGACCAAGACAA ACTGCATTGCCCGCATCTTCGATGGGGTCAAGAATGGCGCTCAGAGGCCAAAATTCATCAGCCCCAG TTCTCAGGTTTTCATCTGCCTGGGTCTGATCCTGGTGCAAATTGTGATGGTGTCTGTGTGGCTCATC CTGGAGGCCCCAGGCACCAGGAGGTATACCCTTACAGAGAAGCGGGAAACAGTCATCCTAAAATGCA ATGTCAAAGATTCCAGCATGTTGATCTCTCTTACCTACGATGTGATCCTGGTGATCTTATGCACTGT GTACGCCTTCAAAACGCGGAAGTGCCCAGAAAATTTCAACGAAGCTAAGTTCATAGGTTTTACCATG TACACCACGTGCATCATCTGGTTGGCCTTCCTCCCTATATTTTATGTGACATCAAGTGACTACAGAC CTCTGCAAGCACGTATGTGTCAACGGTGTGCAATGGGCGGGAAGTCCTCGACTCCACCACCTCATCT CTGTGATTGTGAATTGCAGTTCAGTTCTTGTGTTTTTAGACTGTTAGACAAAAGTGCTCACGTGCAG CTCCAGAATATGGAAACAGAGCAAAAGAACAACCCTAGTACCTTTTTT**TAG**AAACAGTACGATAAAT TATTTTTGAGGACTGTATATAGTGATGTGCTAGAACTTTCTAGGCTGAGTCTAGTGCCCCTATTATT AACAATTCCCCCAGAACATGGAAATAACCATTGTTTACAGAGCTGAGCATTGGTGACAGGGTCTGAC ATGGTCAGTCTACTTCAAG		
ORF Start: ATG at 88		ORF Stop: TAG at 2662

	SEQ ID NO: 168	858 aa	MW at 96975.6kD
NOV40a, CG151014-01 Protein Sequence	MKMLTRLQVLTLALFSKGFLLSLGDHNFLRREIKIEGDLVLGGLFPINEKGTGTEECGRINEDRGIQ RLEAMLFAIDEINKDDYLLPGVKLGVHILDTCSRDTYALEQSLEFVRASLTKVDEAEYMCPDGSYAI QENIPLLIAGVIGGSYSSVSIQVANLLRLFQIPQISYASTSAKLSDKSRYDYFARTVPPDFYQAKAM AEILRFFNWTYVSTVASEGDYGETGIEAFEQEARLRNICIATAEKVGRSNIRKSYDSVIRELLQKPN ARVVVLFMRSDDSRELIAAASRANASFTWVASDGWGAQESIIKGSEHVAYGAITLELASQPVRQFDR YFQSLNPYNNHRNPWFRDFWEQKFQCSLQNKRNHRRVCDKHLAIDSSNYEQESKIMFVVNAVYAMAH ALHKMQRTLCPNTTKLCDAMKILDGKKLYKDYLLKINFTAPFNPNKDADSIVKFDTFGDGMGRYNVF NFQNVGGKYSYLKVGHWAETLSLDVNSIHWSRNSVPTSQCSDPCAPNEMKNMQPGDVCCWICIPCEP YEYLADEFTCMDCGSGQWPTADLTGCYDLPEDYIRWEDAWAIGPVTIACLGFMCTCMVVTVFIKHNN TPLVKASGRELCYILLFGVGLSYCMTFFFIAKPSPVICALRRLGLGSSFAICYSALLTKTNCIARIF DGVKNGAQRPKFISPSSQVFICLGLILVQIVMVSVWLILEAPGTRRYTLTEKRETVILKCNVKDSSM LISLTYDVILVILCTVYAFKTRKCPENFNEAKFIGFTMYTTCIIWLAFLPIFYVTSSDYRPLQARMC QRCAMGGKSSTPPPHLCDCELQFSSCVFRLLDKSAHVQLQNMETEQKNNPSTFF

	SEQ ID NO: 169	1758 bp	
NOV40b, CG151014-02 DNA Sequence	CAAAGATCCAGTTTGGAAATGAGAGAGGACTAGCATGACACATTGGCTCCACCATTGATATCTCCCA GAGGTACAGAAACAGGATTCATGAAGATGTTGACAAGACTGCAAGTTCTTACCTTAGCTTTGTTTTC AAAGGGATTTTTACTCTCTTTAGGGGACCATAACTTTCTAAGGAGAGAGATTAAAATAGAAGGTGAC CTTGTTTTAGGGGGCCTGTTTCCTATTAACGAAAAAGGCACTGGAACTGAAGAATGTGGGCGAATCA ATGAAGACCGAGGGATTCAACGCCTGGAAGCCATGTTGTTTGCTATTGATGAAATCAACAAAGATGA TTACTTGCTACCAGGAGTGAAGTTGGGTGTTCACATTTTGGATACATGTTCAAGGGATACCTATGCA TTGGAGCAATCACTGGAGTTTGTCAGGGCATCTTTGACAAAAGTGGATGAAGCTGAGTATATGTGTC CTGATGGATCCTATGCCATTCAAGAAAACATCCCACTTCTCATTGCAGGGGTCATTGGTGGCTCTTA TAGCAGTGTTTCCATACAGGTGGCAAACCTGCTGCGGCTCTTCCAGATCCCTCAGATCAGCTACGCA TCCACCAGCGCCAAACTCAGTGATAAGTCGCGCTATGATTACTTTGCCAGGACCGTGCCCCCCGACT TCTACCAGGCCAAAGCCATGGCTGAGATCTTGCGCTTCTTCAACTGGACCTACGTGTCCACAGTAGC CTCCGAGGGTGATTACGGGGAGACAGGGATCGAGGCCTTCGAGCAGGAAGCCCGCCTGCGCAACATC TGCATCGCTACGGCGGAGAAGGTGGGCCGCTCCAACATCCGCAAGTCCTACGACAGCGTGATCCGAG AACTGTTGCAGAAGCCCAACGCGCGTGCCTGCTCCTTCATGCGCAGCGACGACTCGCGGGAGCT CATTGCAGCCGCCAGCGCCAATGCCTCCTTCACCTGGGTGGCCAGCGACGGCTGGGGCGCGCAG GAGAGCATCATCAAGGGCAGCGAGCATGTGGCCTACGGCGCCATCACCCTGGAGCTGGCCTCCCAGC

	CTGTCCGCCAGTTCGACCGCTACTTCCAGAGCCTCAACCCCTACAACAACCACCGCAACCCCTGGTT CCGGGACTTCTGGGAGCAAAAGTTTCAGTGCAGCCTCCAGAACAAACGCAACCACAGGCGCGTCTGC GACAAGCACCTGGCCATCGACAGCAGCAACTACGAGCAAGAGTCCAAGATCATGTTTGTGGTGAACG CGGTGTATGCCATGGCCCACGCTTTGCACAAAATGCAGCGCACCCTCTGTCCCAACACTACCAAGCT TTGTGATGCTATGAAGATCCTGGATGGGAAGAAGTTGTACAAGGATTACTTGCTGAAAATCAACTTC ACGGGTGCAGACGACAACCATGTGCATCTCCGTCAGCCTGAGTGGCTTTGTGGTCTTGGGCTGTTTG TTTGCACCCAAGGTTCACATCATCCTGTTTCAACCCCAGAAGAATGTTGTCACACACAGACTGCACC TCAACAGGTTCAGTGTCAGTGGAACTGGGACCACATACTCTCAGTCCTCTGCAAGCACGTATGTGCC AACGGTGTGCAATGGGCGGGAAGTCCTCGACTCCACCACCTCATCTCTGTGATTGTGAATTGCAGTT CAGTTCTTGTGTTTTTAGACTGTT**TAG**ACAAAAGTGCTCACGTGCAGCTCCAGAATATGGAAACAGAG CAAAAGAACAACCCTA		
	ORF Start: ATG at 88		ORF Stop: TAG at 1699

	SEQ ID NO: 170	537 aa	MW at 60801.8kD
NOV40b, CG151014-02 Protein Sequence	MKMLTRLQVLTLALFSKGFLLSLGDHNFLRREIKIEGDLVLGGLFPINEKGTGTEECGRINEDRGIQ RLEAMLFAIDEINKDDYLLPGVKLGVHILDTCSRDTYALEQSLEFVRASLTKVDEAEYMCPDGSYAI QENIPLLIAGVIGGSYSSVSIQVANLLRLFQIPQISYASTSAKLSDKSRYDYFARTVPPDFYQAKAM AEILRFFNWTYVSTVASEGDYGETGIEAFEQEARLRNICIATAEKVGRSNIRKSYDSVIRELLQKPN ARVVVLFMRSDDSRELIAAASRANASFTWVASDGWGAQESIIKGSEHVAYGAITLELASQPVRQFDR YFQSLNPYNNHRNPWFRDFWEQKFQCSLQNKRNHRRVCDKHLAIDSSNYEQESKIMFVVNAVYAMAH ALHKMQRTLCPNTTKLCDAMKILDGKKLYKDYLLKINFTGADDNHVHLRQPEWLCGLGLFVCTQGSH HPVSTPEECCHTQTAPQQVQCQWNWDHILSVLCKHVCANGVQWAGSPRLHHLISVIVNCSSVLVFLD C		

	SEQ ID NO: 171	1758 bp	
NOV40c, CG151014-03 DNA Sequence	CCTTGATCCAGTTTGGAAATGAGAGAGGACTAGCATGACACATTGGCTCCACCATTGATATCTCCCA GAGGTACAGAAACAGGATTC**ATG**AAGATGTTGACAAGACTGCAAGTTCTTACCTTAGCTTTGTTTTC AAAGGGATTTTTACTCTCTTTAGGGGACCATAACTTTCTAAGGAGAGAGATTAAAATAGAAGGTGAC CTTGTTTTAGGGGGCCTGTTTCCTATTAACGAAAAAGGCACTGGAACTGAAGAATGTGGGCGAATCA ATGAAGACCGAGGGATTCAACGCCTGGAAGCCATGTTGTTTGCTATTGATGAAATCAACAAAGATGA TTACTTGCTACCAGGAGTGAAGTTGGGTGTTCACATTTTGGATACATGTTCAAGGGATACCTATGCA TTGGAGCAATCACTGGAGTTTGTCAGGGCATCTTTGACAAAAGTGGATGAAGCTGAGTATATGTGTC CTGATGGATCCTATGCCATTCAAGAAAACATCCCACTTCTCATTGCAGGGGTCATTGGTGGCTCTTA TAGCAGTGTTTCCATACAGGTGGCAAACCTGCTGCGGCTCTTCCAGATCCCTCAGATCAGCTACGCA TCCACCAGCGCCAAACTCAGTGATAAGTCGCGCTATGATTACTTTGCCAGGACCGTGCCCCCCGACT TCTACCAGGCCAAAGCCATGGCTGAGATCTTGCGCTTCTTCAACTGGACCTACGTGTCCACAGTAGC CTCCGAGGGTGATTACGGGGAGACAGGGATCGAGGCCTTCGAGCAGGAAGCCCGCCTGCGCAACATC TGCATCGCTACGGCCGGAGAAGGTGGGCCGCTCCAACATCCGCAAGTCCTACGACAGCGTGATCCGAG AACTGTTGCAGAAGCCCAACGCGCGCGTCGTGGTCCTCTTCATGCGCAGCGACGACTCGCGGGAGCT CATTGCAGCCGCCAGCCGCGCCAATGCCTCCTTCACCTGGGTGGCCAGCGACGGCTGGGGCGCGCAG GAGAGCATCATCAAGGGCAGCGAGCATGTGGCCTACGGCGCCATCACCCTGGAGCTGGCCTCCCAGC CTGTCCGCCAGTTCGACCGCTACTTCCAGAGCCTCAACCCCTACAACAACCACCGCAACCCCTGGTT CCGGGACTTCTGGGAGCAAAAGTTTCAGTGCAGCCTCCAGAACAAACGCAACCACAGGCGCGTCTGC GACAAGCACCTGGCCATCGACAGCAGCAACTACGAGCAAGAGTCCAAGATCATGTTTGTGGTGAACG CGGTGTATGCCATGGCCCACGCTTTGCACAAAATGCAGCGCACCCTCTGTCCCAACACTACCAAGCT TTGTGATGCTATGAAGATCCTGGATGGGAAGAAGTTGTACAAGGATTACTTGCTGAAAATCAACTTC ACGGGTGCAGACGACAACCATGTGCATCTCCGTCAGCCTGAGTGGCTTTGTGGTCTTGGGCTGTTTG TTTGCACCCAAGGTTCACATCATCCTGTTTCAACCCCAGAAGAATGTTGTCACACACAGACTGCACC TCAACAGGTTCAGTGTCAGTGGAACTGGGACCACATACTCTCAGTCCTCTGCAAGCACGTATGTGCC AACGGTGTGCAATGGGCGGGAAGTCCTCGACTCCACCACCTCATCTCTGTGATTGTGAATTGCAGTT CAGTTCTTGTGTTTTTAGACTGTT**TAG**ACAAAAGTGCTCACGTGCAGCTCCAGAATATGGAAACAGAG CAAAAGAACAACCCTA		
	ORF Start: ATG at 88		ORF Stop: TAG at 1699

	SEQ ID NO: 172	537 aa	MW at 60801.8kD
NOV40c, CG151014-03 Protein Sequence	MKMLTRLQVLTLALFSKGFLLSLGDHNFLRREIKIEGDLVLGGLFPINEKGTGTEECGRINEDRGIQ RLEAMLFAIDEINKDDYLLPGVKLGVHILDTCSRDTYALEQSLEFVRASLTKVDEAEYMCPDGSYAI QENIPLLIAGVIGGSYSSVSIQVANLLRLFQIPQISYASTSAKLSDKSRYDYFARTVPPDFYQAKAM AEILRFFNWTYVSTVASEGDYGETGIEAFEQEARLRNICIATAEKVGRSNIRKSYDSVIRELLQKPN ARVVVLFMRSDDSRELIAAASRANASFTWVASDGWGAQESIIKGSEHVAYGAITLELASQPVRQFDR YFQSLNPYNNHRNPWFRDFWEQKFQCSLQNKRNHRRVCDKHLAIDSSNYEQESKIMFVVNAVYAMAH ALHKMQRTLCPNTTKLCDAMKILDGKKLYKDYLLKINFTGADDNHVHLRQPEWLCGLGLFVCTQGSH HPVSTPEECCHTQTAPQQVQCQWNWDHILSVLCKHVCANGVQWAGSPRLHHLISVIVNCSSVLVFLD C		

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Sequence comparison of the above protein sequences yields the following sequence relationships shown in Table 40B.

Table 40B. Comparison of NOV40a against NOV40b and NOV40c.		
Protein Sequence	NOV40a Residues/ Match Residues	Identities/ Similarities for the Matched Region
NOV40b	1..441 1..441	409/441 (92%) 409/441 (92%)
NOV40c	1..441 1..441	409/441 (92%) 409/441 (92%)

10

Further analysis of the NOV40a protein yielded the following properties shown in Table 40C.

15

Table 40C. Protein Sequence Properties NOV40a	
PSort analysis:	0.6400 probability located in plasma membrane; 0.4600 probability located in Golgi body; 0.3700 probability located in endoplasmic reticulum (membrane); 0.1000 probability located in endoplasmic reticulum (lumen)
SignalP analysis:	Cleavage site between residues 25 and 26

A search of the NOV40a protein against the Geneseq database, a proprietary database that contains sequences published in patents and patent publication, yielded

20   several homologous proteins shown in Table 40D.

Table 40D. Geneseq Results for NOV40a

Geneseq Identifier	Protein/Organism/Length [Patent #, Date]	NOV40a Residues/ Match Residues	Identities/ Similarities for the Matched Region	Expect Value
AAE15990	Human glutamate receptor, metabotrophic 3 (GRM3) protein - Homo sapiens, 877 aa. [WO200196350-A2, 20-DEC-2001]	3..811 1..809	797/809 (98%) 799/809 (98%)	0.0
AAR82657	Human mGluR3 - Homo sapiens, 877 aa. [WO9522609-A2, 24-AUG-1995]	3..811 1..809	797/809 (98%) 799/809 (98%)	0.0
AAM23698	Human EST encoded protein SEQ ID NO: 1223 - Homo sapiens, 857 aa. [WO200154477-A2, 02-AUG-2001]	1..811 1..811	796/811 (98%) 798/811 (98%)	0.0
AAR64252	Human mGluR3 - Homo sapiens, 879 aa. [WO9429449-A, 22-DEC-1994]	1..811 1..811	796/811 (98%) 799/811 (98%)	0.0
AAO15105	Human ph2SPMGluR3-CaR*AAA* Gqi5 fusion construct protein sequence - Chimeric - Homo sapiens, 1402 aa. [WO200229033-A2, 11-APR-2002]	21..811 17..807	777/791 (98%) 781/791 (98%)	0.0

In a BLAST search of public sequence datbases, the NOV40a protein was found to have homology to the proteins shown in the BLASTP data in Table 40E.

5

Table 40E. Public BLASTP Results for NOV40a				
Protein Accession Number	Protein/Organism/Length	NOV40a Residues/ Match Residues	Identities/ Similarities for the Matched Portion	Expect Value
Q14832	Metabotropic glutamate receptor 3 precursor (mGluR3) - Homo sapiens (Human), 877 aa.	3..811 1..809	797/809 (98%) 799/809 (98%)	0.0

Q8TBH9	Glutamate receptor, metabotropic 3 – Homo sapiens (Human), 877 aa.	3..811 1..809	795/809 (98%) 797/809 (98%)	0.0
Q9QYS2	Metabotropic glutamate receptor 3 protein – Mus musculus (Mouse), 879 aa.	1..811 1..811	773/811 (95%) 792/811 (97%)	0.0
P31422	Metabotropic glutamate receptor 3 precursor – Rattus norvegicus (Rat), 879 aa.	1..811 1..811	772/811 (95%) 790/811 (97%)	0.0
JC7160	metabotropic glutamate receptor subtype 3 precursor – mouse, 879 aa.	1..811 1..811	771/811 (95%) 790/811 (97%)	0.0

PFam analysis predicts that the NOV40a protein contains the domains shown in the Table 40F.

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Table 40F. Domain Analysis of NOV40a			
Pfam Domain	NOV40a Match Region	Identities/ Similarities for the Matched Region	Expect Value
ANF_receptor	58..489	194/473 (41%) 399/473 (84%)	3.2e-173
7tm_3	576..820	109/283 (39%) 217/283 (77%)	3.1e-104

Example 41.

10   The NOV41 clone was analyzed, and the nucleotide and encoded polypeptide sequences are shown in Table 41A.

Table 41A. NOV41 Sequence Analysis			
	SEQ ID NO: 173	880 bp	
NOV41a, CG151297-01 DNA Sequence	GAATTCTGATGTGCTTCAGTGCACAGAACAGTAACAGATGAGCTGCTTTTGGGGAGAGCTTGAGTAC TCAGTCGGAGCATCATCATGGGGTCTAGTGCCACAGAGATTGAAGAATTGGAAAACACCACTTTTAA GTATCTTACAGGAGAACAGACTGAAAAAATGTGGCAGCGCCTGAAAGGAATACTAAGATGCTTGGTG AAGCAGCTGGAAAGAGGTGATGTTAACGTCGTCGACTTAAAGAAGAATATTGAATATGCGGCATCTG TGCTGGAAGCAGTTTATATCGATGAAACAAGAAGACTTCTGGATACTGAAGATGAGCTCAGTGACAT TCAGACTGACTCAGTCCCATCTGAAGTCCGGGACTGGTTGGCTTCTACCTTTACACGGAAAATGGGG ATGACAAAAAGAAACCTGAGGAAAAACCAAAATTTCGGAGCATTGTGCATGCTGTTCAAGCTGGAA TTTTTGTGGAAAGAATGTACCGAAAAACATTTTCTCTTCTGACAGACTCAACAGAGAAAATTGTTAT TCCTCTTATAGAGGAAGCCTCAAAAGCCGAAACTTCTTCCTATGTGGCAAGCAGCTCAACCACCATT GTGGGGTTACACATTGCTGATGCACTAAGACGATCAAATACAAAAGGCTCCATGAGTGATGGGTCCT ATTCCCCAGACTACTCCCTTGCAGCAGTGGACCTGAAGAGTTTCAAGAACAACCTGGTGGACATCAT TCAGCAGAACAAAGAGAGGTGGAAAGAGTTAGCTGCACAAGAAGCAAGAACCAGTTCACAGAAGTGT

	GAGTTTATTCATCAGTA**AA**CACCTTTAAGTAAAAC~~GTCGTGCATGGTGGCAGCTC~~TAATTGACCAA AAGACTTGG		
	ORF Start: ATG at 85		ORF Stop: TAA at 820

	SEQ ID NO: 174	245 aa	MW at 27787.2kD
NOV41a, CG151297-01 Protein Sequence	MGSSATEIEELENTTFKYLTGEQTEKMWQRLKGILRCLVKQLERGDVNVVDLKKNIEYAASVLEAVY IDETRRLLDTEDELSDIQTDSVPSEVRDWLASTFTRKMGMTKKKPEEKPKFRSIVHAVQAGIFVERM YRKTFSLLTDSTEKIVIPLIEEASKAETSSYVASSSTTIVGLHIADALRRSNTKGSMSDGSYSPDYS LAAVDLKSFKNNLVDIIQQNKERWKELAAQEARTSSQKCEFIHQ		

	SEQ ID NO: 175	1817 bp	
NOV41b, CG151297-02 DNA Sequence	<u>TCAGTGCACAGAACAGTAACAGATGAGCTGCTTTTGGGGAGAGCTTGAGTACTCAGTCGGTCAGTAG</u> <u>TACAGTAGCAGGCTCACATGTACGGATTGTTCTTGTGAGGAGCATCATC</u>**ATG**GGGTCTAGTGCCACA GAGATTGAAGAATTGGAAAACACCCACTTTTAAGTATCTTACAGGAGAACAGACTGAAAAAATGTGGC AGCGCCTGAAAGGAATACTAAGATGCTTGGTGAAGCAGCTGGAAAGAGGTGATGTTAACGTCGTCGA CTTAAAGAAGAATATTGAATATGCGGCATCTGTGCTGGAAGCAGTTTATATCGATGAAACAAGAAGA CTTCTGGATACTGAAGATGAGCTCAGTGACATTCAGACTGACTCAGTCCCATCTGAAGTCCGGGACT GGTTGGCTTCTACCTTTACACGGAAAATGGGGATGACAAAAAAGAAACCTGAGGAAAAACCAAAATT TCGGAGCATTGTGCATGCTGTTCAAGCTGGAATTTTTGTGGAAGAATGTACCGAAAAACATATCAT ATGGTTGGTTTGGCATATCCAGCAGCTGTCATCGTAACATTAAAGGATGTTGATAAATGGTCTTTCG ATGTATTTGCCCTAAATGAAGCAAGTGGAGAGCATAGTCTGAAGTTTATGATTTATGAACTGTTTAC CAGATATGATCTTATCAACCGTTTCAAGATTCCTGTTTCTTGCCTAATCACCTTTGCAGAAGCTTTA GAAGTTGGTTACGGCAAGTACAAAAATCCATATCACAATTTGATTCATGCAGCTGATGTCACTCAAA CTGTGCATTACATAATGCTTCATACAGGTATCATGCACTGGCTCACTGAACTGGAAATTTTTAGCAAT GGTCTTTGCTGCTGCCATTCATGATTATGAGCATACAGGGACAACAAACAACTTTCACATTCAGACA AGGTCAGATGTTGCCATTTTTGTATAATGATCGCTCTGTCCTTGAGAATCACCACGTGAGTGCAGCTT ATCGACTTATGCAAGAAGAAGAAATGAATATCTTGATAAATTTATCCAAAGATGACTGGAGGGATCT TCGGAACCTAGTGATTGAAATGGTTTTATCTACAGACATGTCAGGTCACTTCCAGCAAATTAAAAAT ATAAGAAACAGTTTGCAGCAGCCTGAAGGGATTGACAGAGCCAAAACCATGTCCCTGATTCTCCACG CAGCAGACATCAGCCACCCAGCCAAATCCTGGAAGCTGCATTATCGGTGGACCATGGCCCTAATGGA GGAGTTTTTCCTGCAGGGAGATAAAGAAGCTGAATTAGGGCTTCCATTTTCCCCACTTTGTGATCGG AAGTCAACCATGGTGGCCCAGTCACAAATAGGTTTCATCGATTTCATAGTAGAGCCAACATTTTCTC TTCTGACAGACTCAACAGAGAAAATTGTTATTCCTCTTATAGAGGAAGCCTCAAAAGCCGAAACTTC TTCCTATGTGGCAAGCAGCTCAACCACCATTGTGGGGTTACACATTGCTGATGCACTAAGACGATCA AATACAAAAGGCTCCATGAGTGATGGGTCCTATTCCCCAGACTACTCCCTTGCAGCAGTGGACCTGA AGAGTTTCAAGAACAACCTGGTGGACATCATTCAGCAGACAAAGAGAGGTGGAAAGAGTTAGTTGC ACAAGAAGCAAGAACCAGTTCACAGAAGTGTGAGTTTATTCATCAGTA**AA**<u>CACCTTTAAGTAAAACC</u> <u>TCGTGCATGGTGGCAGCTCTAATTTGACCAAAAGACTTGGAGATTTTGATTATGCTTGCTGGATATC</u> <u>TATTCTGT</u>		
	ORF Start: ATG at 117		ORF Stop: TAA at 1722

	SEQ ID NO: 176	535 aa	MW at 61249.3kD
NOV41b, CG151297-02 Protein Sequence	MGSSATEIEELENTTFKYLTGEQTEKMWQRLKGILRCLVKQLERGDVNVVDLKKNIEYAASVLEAVY IDETRRLLDTEDELSDIQTDSVPSEVRDWLASTFTRKMGMTKKKPEEKPKFRSIVHAVQAGIFVERM YRKTYHMVGLAYPAAVIVTLKDVDKWSFDVFALNEASGEHSLKFMIYELPTRYDLINRFKIPVSCLI TFAEALEVGYGKYKNPYHNLIHAADVTQTVHYIMLHTGIMHWLTELEILAMVFAAAIHDYEHTGTTN NFHIQTRSDVAILYNDRSVLENHHVSAAYRLMQEEEMNILINLSKDDWRDLRNLVIEMVLSTDMSGH FQQIKNIRNSLQQPEGIDRAKTMSLILHAADISHPAKSWKLHYRWTMALMEEFFLQGDKEAELGLPF SPLCDRKSTMVAQSQIGFIDFIVEPTFSLLTDSTEKIVIPLIEEASKAETSSYVASSSTTIVGLHIA DALRRSNTKGSMSDGSYSPDYSLAAVDLKSFKNNLVDIIQQNKERWKELVAQEARTSSQKCEFIHQ		

Sequence comparison of the above protein sequences yields the following sequence relationships shown in Table 41B.

Table 41B. Comparison of NOV41a against NOV41b.		
Protein Sequence	NOV41a Residues/ Match Residues	Identities/ Similarities for the Matched Region
NOV41b	1..159 1..159	141/159 (88%) 148/159 (92%)

Further analysis of the NOV41a protein yielded the following properties shown in Table 41C.

Table 41C. Protein Sequence Properties NOV41a	
PSort analysis:	0.8800 probability located in nucleus; 0.1000 probability located in mitochondrial matrix space; 0.1000 probability located in lysosome (lumen); 0.1000 probability located in plasma membrane
SignalP analysis:	No Known Signal Sequence Predicted

A search of the NOV41a protein against the Geneseq database, a proprietary database that contains sequences published in patents and patent publication, yielded several homologous proteins shown in Table 41D.

Table 41D. Geneseq Results for NOV41a				
Geneseq Identifier	Protein/Organism/Length [Patent #, Date]	NOV41a Residues/ Match Residues	Identities/ Similarities for the Matched Region	Expect Value
AAB85116	Human 3', 5' cyclic nucleotide phosphodiesterase (HSPDE1A3A) - Homo sapiens, 535 aa. [EP1097707-A1, 09-MAY-2001]	1..159 1..159	141/159 (88%) 148/159 (92%)	5e-75
AAB85105	Human 3', 5' cyclic nucleotide phosphodiesterase (HSPDE1A3A) - Homo sapiens. 535 aa.	1..159 1..159	141/159 (88%) 148/159 (92%)	5e-75

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	[EP1097706-A1, 09-MAY-2001]			
AAE07953	Human phosphodiesterase (PDE) type 1 protein - Homo sapiens, 535 aa. [EP1097719-A1, 09-MAY-2001]	1..159 1..159	141/159 (88%) 148/159 (92%)	5e-75
AAE07917	Human phosphodiesterase (PDE) type 1 protein - Homo sapiens, 535 aa. [EP1097718-A1, 09-MAY-2001]	1..159 1..159	141/159 (88%) 148/159 (92%)	5e-75
AAY80988	Human 61 kD CaM-PDE (clone pHcam61-6N-7), SEQ ID NO:49 - Homo sapiens, 535 aa. [US6015677-A, 18-JAN-2000]	1..159 1..159	141/159 (88%) 148/159 (92%)	5e-75

In a BLAST search of public sequence datbases, the NOV41a protein was found to have homology to the proteins shown in the BLASTP data in Table 41E.

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Table 41E. Public BLASTP Results for NOV41a				
Protein Accession Number	Protein/Organism/Length	NOV41a Residues/ Match Residues	Identities/ Similarities for the Matched Portion	Expect Value
AAH22480	Hypothetical 62.3 kDa protein - Homo sapiens (Human), 545 aa.	1..159 1..159	141/159 (88%) 148/159 (92%)	1e-74
P54750	Calcium/calmodulin-dependent 3',5'-cyclic nucleotide phosphodiesterase 1A (EC 3.1.4.17) (Cam-PDE 1A) (61 kDa Cam-PDE) (hCam-1) - Homo sapiens (Human), 534 aa.	2..159 1..158	140/158 (88%) 147/158 (92%)	6e-74
Q9EPR9	Phosphodiesterase 1A - Rattus norvegicus (Rat), 542 aa.	1..159 1..159	134/159 (84%) 144/159 (90%)	6e-71
Q61481	Calcium/calmodulin-dependent 3',5'-cyclic nucleotide phosphodiesterase 1A (EC 3.1.4.17) (Cam-PDE 1A) (61 kDa Cam-PDE) - Mus musculus (Mouse), 565 aa.	1..159 21..179	133/159 (83%) 143/159 (89%)	3e-70

A45334	3',5'-cyclic-nucleotide phosphodiesterase (EC 3.1.4.17) 1A, calmodulin-dependent, 61K brain form - bovine, 530 aa.	1..159 1..159	129/159 (81%) 144/159 (90%)	6e-69

PFam analysis predicts that the NOV41a protein contains the domains shown in the Table 41F.

Table 41F. Domain Analysis of NOV41a

Pfam Domain	NOV41a Match Region	Identities/ Similarities for the Matched Region	Expect Value
PDEase	138..159	9/49 (18%) 22/49 (45%)	0.11

Example 42.

The NOV42 clone was analyzed, and the nucleotide and encoded polypeptide sequences are shown in Table 42A.

Table 42A. NOV42 Sequence Analysis

	SEQ ID NO: 177	512 bp	
NOV42a, CG151822-01 DNA Sequence	CCATGGCGGGCTGCGCGGCGCGGGCTCCGCCGGGCTCTGAGGCGCGTCTCAGCCTCGCCACCTTCCT GCTGGGCGCCTCGGTGCTCGCGCTGCCGCTGCTCACGCGCGCCGGCCTGCAGGGCCGCACCGGGCTG GCGCTCTACGTGGCCGGGCTCAACGCGCTGCTGCTGCTGCTCTATCGGCCGCCTCGCTACCAGATAG CCATCCGAGCTTGTTTCCTGGGGTTTGTGTTCGGCTGCGGCACGCTGCTAAGTTTTAGCCAGTCTTC TTGGAGTCACTTTGGCT**GA**ACTGAAGCAGATTACCTGGCTCAGTGTCACAGGGCTGCTGATGGTGGT CTTCGGAGAATGTCTGAGGAAGGCGGCCATGTNTACAGCTGGCTCCAATTTCAACCACGTGGTACAG ACGTCGGGTGGTTTTACTGGAGTATTGGAACTCAGGTGATGCT		
	ORF Start: ATG at 3		ORF Stop: TGA at 285

	SEQ ID NO: 178	94 aa	MW at 9871.5kD
NOV42a, CG151822-01 Protein Sequence	MAGCAARAPPGSEARLSLATFLLGASVLALPLLTRAGLQGRTGLALYVAGLNALLLLLYRPPRYQIA IRACFLGFVFGCGTLLSFSQSSWSHFG		

	SEQ ID NO: 179	3597 bp	
NOV42b, CG151822-02 DNA Sequence	GGCACGAGCGGCGCCGCCGCCCGCTAGTCCGCCGCCCGGCGCC**ATG**GCGGGCTGCGCGGCGCGGGCT CCGCCGGGGCTCTGAGGCGCGTCTCAGCCTCGCCACCTTCCTGCTGGGCGCCTCGGTGCTCGCGCTGC CGCTGCTCACGCGCGCCGGCCTGCAGGGCCGCACCGGGCTGGCGCTCTACGTGGCCGGGCTCAACGC GCTGCTGCTGCTGCTCTATCGGCCGCCTCGCTACCAGATAGCCATCCGAGCTTGTTTCCTGGGGGTTT GTGTTCGGCTGCGGCACGCTGCTAAGTTTTAGCCAGTCTTCTTGGAGTCACTTTGGCTGGTACATGT GCTCCCTGTCATTGTTCCACTATTCTGAATACTTGGTGACAGCAGTCAATAATCCCAAAAGTCTGTC CTTGGATTCCTTTCTCCTGAATCACAGCCTGGAGTATACAGTAGCTGCTCTTTCTTCTTGGTTAGAG TTCACACTTGAAAATATCTTTTGGCCAGAACTGAAGCAGATTACCTGGCTCAGTGTCACAGGGCTGC TGATGGTGGTCTTCGGAGAATGTCTGAGGAAGGCGGCCATGTTTACAGCTGGCTCCAATTTCAACCA CGTGGTACAGAATGAAAAATCAGATACACATACTCTGGTGACCAGTGGAGTGTACGCTTGGTTTCGG CATCCTTCTTACGTCGGGTGGTTTTACTGGAGTATTGGAACTCAGGTGATGCTGTGTAACCCCATCT GCGGCGTCAGCTATGCCCTGACAGTGTGGCGATTCTTCCGCGATCGAACAGAAGAAGAAGAAATCTC ACTAATTCACTTTTTTGGAGAGGAGTACCTGGAGTATAAGAAGAGGGTGCCCACGGGCCTGCCTTTC ATAAAGGGGGTCAAGGTGGACCTGTGA<u>CGGGCAGTGGCCCCGGTGACCTTGGGGCCTCCGACCCTGT GCAGCCTGGGACAAAACTGTTTCCGGTTGGCCGCTGCCACATGGATTTTCTTAATCGTTTTATGTCA TTAGTCACTCTTCTGGAATGTCACTCAAGACCAAGCGGTCAGAAGGCCTGAGGACCCAAGGCCCCAC TGGAGCAGTCTGTCCTTATGCCGAATCAAGGCGGAACATGGGTGAAAGACGAGTAAGGGGCAAATCA CAGCAATATTCCACAGCGCCTCCAGAGTTACCTGGGGAGGACCGAGGCCACACGCCACTGCCCCCG AGGCCAGAGTGTAAGTAAAGGATAACCAGGACTCGCTGGGAGAGATGGACTCTGTCCTCAGCAACAC TCCACAGCAGAAAGGGGTAGCAGGTACCCCTTCTTATCAGCGGTAAAAATGCATTTACAACCTTTCA TTTAACCGAAAAACACAGACCGCTTTAACCTCTTTATTTCTGTCCCCACTGCATGAACATCTATAC AATTTTAAAAATACTTCCTCATAGGATGGCTTTGGCCCTTCATCTATTTAATCATAGCTACATACCTA TTTTTTTATAAGTACGCAGTACACATTCAAAGGGGTATTCCTAGCTCAATGCTTGGTGTTCTAGTTCA ACTTTTATCCTGCAGCAAGTAAGCCTAGATAACTCTACACGATTTGGCTGAGTGGCTTTGTGTGACC GTGGCCCCAGGCCAAGGGGACCATGGCCCTGGCTGGCTTTCCCCCGGGGGTCTCAGCTCCTGTTGTC AGTGATAGGCGGCTCAAAGGAGCATCAGTTTCTTTTGATCCAAGAAGTGCTTACTGAATGCCTGCCC TGTGCGTGGCCTTAAACATTGAGAAGTGCTGCTCTCCGTTTATTTGGGATTTGATTCTCATTTTACC ATAGCTTATATTCTCAATTTCAATGCCAGTCTCAGAACTCTTGTTTTCTGTGTTCTGTTCTCAAAAT TACATTGTCCCTCATGTCATTTCAAACTGTTTTCCAAAGGGATTTGAGCATATACAACTACAAATCC AAGCAGATTGACTCTCAAAAATAATCTTAAATACTGCAAATAGTCCCAACTAAGATTCAGTCAGTAT GTTTGTTTTGCAAGTTTTGGGAGGTAAGTTGGCTTTGAGTCACACATCGAAGCTTTAAGAGGTGAGA CGCTGGCTTCATTCTGGACTAGACAGGAACTTGGCCTCAGCGTGAGATCCTGCCATGCAGTGTTGCG GTGGCACTGAAGAAGTGTGAATGTGAAGGCGGCGTCGGCGCGGGGCCAGAGCACCACTCTGCTGCCC CACCACGCGGCCTGTGAGGAGCCACTAAACCTTTCCGTGCCTAGACCTCCCCATCTGTGGAATGGGG TCAATACCACCTACCTCACCAGGGGTGTTGTGAGGAACAATGTCAAATGTTTTTAATAC TCAGATGTGGGAGCGACATCAATGAAATCTGTACTGTATGAAAGCTACACAAAAATGGGCAGACATT TGGTTAATTGTGCCAGATACCTAAAATGTATGTTCAGAAAAGCATTTTATCAACTCAGAAATATGAC TTATTTCTAGATTCATGGCTTAATGAATTTTTTTCATTGTTATATATACCAAAGAGGCTTACGGGTTC ATTGATTGGTTTGAAAACCAGACAGACGGCCGGGCACGCCTGTAATCCCAAAGTGCTGGGATTGCAG CGTGAGCCACCACGCCCAGCCAAGATGAACTCCTTAAGGACAGGATTTGGTAAGTGATTGACTTCTT TTTAGTTCCATGATCTTGAGATTATTTTTAGCTTTATAAATTTAGCAGTGGCAGGGCCCGTGGGAGAA TCAGGTTAATGAGGTAAAGGCTTTCTGGGTATTTGCTGCCAAGGCCACATCACCAATTTTTCTCGATT TAAAAAACTGTCAAGAGATTTTATTTTTTCCATTGCAGGTGTTTAAAGTGGAGATTCTGAAGTGGAAAAT AGGTACTGTCAGAACAAAGCTACCTGGAAACAGCATAGAGTGAAGCCTTTCGTGAGGGCTTGCAGGC CGCTGCTGAGTGGCAGTTTACAGAAGAGGTCGCGGGGTGAGCCTCTTAGCAGGACAGAAAACAAGGC AGCAGCGCACCTGCCACCCCTTCACGAGCTGCTCCTTGAGCCTAAAAAGTAGGCTTTATTCATCCCT TCTGTTCATTTACCAACCTGGGGGATTGATACGACGCGGGGAAAATGTTCCTAAACCAGGAAGCTGCG TTAGCCGATCAGGCTTTGTAAGATCTCGCCAACAGCTAGCTGCTTAGGAGTACCCCCACGATACGCA CAGCACACCACTGTCCCTTCACTGCACTTTCTTCCTGCCTTAGGTAGTTGGGCTTGCCCACCCTAGT TTGCTTTTGTAGTGGTTTGGCAAGGTTAGAAGGCCTCGGCCTCTCTGTCATGCTGGGAAGTGCCTAC TCTCTGGGCCACTGCTGCAGAGGCCGTGGCACTTGTCATGGGTTTGGAAGACCCAGCCATCTGCAGC AGAGGCAGCCTATCCCATTGCAAGGAGAGGAACTGAACGGAGTAATTATTCTACTCTTCTTTTTACA TAAATGTTTATTTAAATATTCTAAATTGGATTTTCATTCACAGATACTGATTATTCTTTCCAGTTCT TAAATAAAACTGCACTTGATTTCACTCAAAAAAAAAAAAAAAAA</u>
	ORF Start: ATG at 44		ORF Stop: TGA at 896

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	SEQ ID NO: 180	284 aa	MW at 31937.7kD
NOV42b, CG151822-02 Protein Sequence	MAGCAARAPPGSEARLSLATFLLGASVLALPLLTRAGLQGRTGLALYVAGLNALLLLLYRPPRYQIA IRACFLGFVFGCGTLLSFSQSSWSHFGWYMCSLSLFHYSEYLVTAVNNPKSLSLDSFLLNHSLEYTV AALSSWLEFTLENIFWPELKQITWLSVTGLLMVVFGECLRKAAMFTAGSNFNHVVQNEKSDTHTLVT SGVYAWFRHPSYVGWFYWSIGTQVMLCNPICGVSYALTVWRFFRDRTEEEEISLIHFFGEEYLEYKK RVPTGLPFIKGVKVDL		

Sequence comparison of the above protein sequences yields the following sequence relationships shown in Table 42B.

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Table 42B. Comparison of NOV42a against NOV42b.		
Protein Sequence	NOV42a Residues/ Match Residues	Identities/ Similarities for the Matched Region
NOV42b	1..94 1..94	67/94 (71%) 67/94 (71%)

Further analysis of the NOV42a protein yielded the following properties shown in Table 42C.

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Table 42C. Protein Sequence Properties NOV42a	
PSort analysis:	0.6000 probability located in plasma membrane; 0.4000 probability located in Golgi body; 0.3174 probability located in mitochondrial intermembrane space; 0.3000 probability located in endoplasmic reticulum (membrane)
SignalP analysis:	Cleavage site between residues 37 and 38

A search of the NOV42a protein against the Geneseq database, a proprietary database that contains sequences published in patents and patent publication, yielded several homologous proteins shown in Table 42D.

Table 42D. Geneseq Results for NOV42a				
Geneseq Identifier	Protein/Organism/Length [Patent #, Date]	NOV42a Residues/ Match Residues	Identities/ Similarities for the Matched Region	Expect Value
AAY32299	Farnesyl-directed cysteine carboxymethyltransferase STE14 - Homo sapiens, 284 aa. [WO9955878-A1, 04-NOV-1999]	1..94 1..94	94/94 (100%) 94/94 (100%)	2e-48
AAW67730	Human prenylcysteine carboxyl methyltransferase - Homo sapiens, 284 aa.	1..94 1..94	94/94 (100%) 94/94 (100%)	2e-48

	[WO9856924-A1, 17-DEC-1998]			
AAB32052	Human secreted protein BLAST search protein SEQ ID NO: 110 - Homo sapiens, 223 aa. [WO200058350-A1, 05-OCT-2000]	12..94 1..83	83/83 (100%) 83/83 (100%)	3e-41
AAB32051	Human secreted protein BLAST search protein SEQ ID NO: 109 - Homo sapiens, 223 aa. [WO200058350-A1, 05-OCT-2000]	12..94 1..83	83/83 (100%) 83/83 (100%)	3e-41
AAY32300	Mouse farnesyl-directed cysteine carboxymethyltransferase - Mus musculus, 153 aa. [WO9955878-A1, 04-NOV-1999]	5..94 4..93	82/90 (91%) 83/90 (92%)	2e-40

In a BLAST search of public sequence datbases, the NOV42a protein was found to have homology to the proteins shown in the BLASTP data in Table 42E.

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Table 42E. Public BLASTP Results for NOV42a				
Protein Accession Number	Protein/Organism/Length	NOV42a Residues/ Match Residues	Identities/ Similarities for the Matched Portion	Expect Value
O60725	Protein-S isoprenylcysteine O-methyltransferase (EC 2.1.1.100) (Isoprenylcysteine carboxylmethyltransferase) (Prenylcysteine carboxyl methyltransferase) (pcCMT) (Prenylated protein carboxyl methyltransferase) (PPMT) - Homo sapiens (Human), 284 aa.	1..94 1..94	94/94 (100%) 94/94 (100%)	5e-48

Q9EQK7	Protein-S isoprenylcysteine O-methyltransferase (EC 2.1.1.100) (Isoprenylcysteine carboxylmethyltransferase) (Prenylcysteine carboxyl methyltransferase) (pcCMT) (Prenylated protein carboxyl methyltransferase) (PPMT) – Mus musculus (Mouse), 283 aa.	5..94 4..93	84/90 (93%) 85/90 (94%)	2e-41
O12947	Protein-S isoprenylcysteine O-methyltransferase (EC 2.1.1.100) (Isoprenylcysteine carboxylmethyltransferase) (Prenylcysteine carboxyl methyltransferase) (pcCMT) (Prenylated protein carboxyl methyltransferase) (PPMT) (Farnesyl cysteine carboxyl methyltransferase) (FCMT) – Xenopus laevis (African clawed frog), 288 aa.	13..94 9..98	49/90 (54%) 59/90 (65%)	2e-19
Q9WVM4	Protein-S isoprenylcysteine O-methyltransferase (EC 2.1.1.100) (Isoprenylcysteine carboxylmethyltransferase) (Prenylcysteine carboxyl methyltransferase) (pcCMT) (Prenylated protein carboxyl methyltransferase) (PPMT) (Farnesyl cysteine carboxyl methyltransferase) (FCMT) – Rattus norvegicus (Rat), 232 aa (fragment).	53..94 1..42	39/42 (92%) 40/42 (94%)	8e-17
Q9R1L8	Farnesyl cysteine carboxyl methyltransferase - Rattus norvegicus (Rat), 33 aa (fragment).	65..94 1..30	28/30 (93%) 29/30 (96%)	4e-10

Example 43.

The NOV43 clone was analyzed, and the nucleotide and encoded polypeptide sequences are shown in Table 43A.

Table 43A. NOV43 Sequence Analysis			
	SEQ ID NO: 181	2306 bp	
NOV43a,	GCCATGGCGTCCTGCGTGGGGAGCCGGACCCTAAGCAAGGATGATGTGAACTACAAAATGCATTTCC		

CG152256-01 DNA Sequence	GGATGATCAACGAGCAGCAAGTGGAGGACATCACCATTGACTTCTTCTACCGGCGGCATAGCATGAG
	CCTGCTCAGCTTCACCATCGTCAGCCTCATGTACTTCGCCTTTACCAGGGATGACTCTGTTCCAGAA
	GACAACATCTGGAGAGGCATCCTCTCTGTTATTTTTCTTCTTTCTTATCATCAGTGTGTTAGCTTTCC
	CCAATGGTCCGTTCACTCGACCTCATCCAGCCTTATGGCGAATGGTTTTTGGACTCAGTGTGCTCTA
	CTTCCTGTTCCTGGTATTCCTACTCTTCCTGAATTTCGAGCAGGTTAAATCTCTAATGTATTGGCTA
	GATCCAAATCTTCGATACGCCACAAGGGAAGCAGATGTCATGGAGTATGCTGTGAACTGCCATGTGA
	TCACCTGGGAGAGGATTATCAGCCACTTTGATATTTTTGCATTTGGACATTTCTGGGGCTGGGCCAT
	GAAGGCCTTGCTGATCCGTAGTTACGGTCTCTGCTGGACAATCAGTATTACCTGGGAGCTGACTGAG
	CTCTTCTTCATGCATCTCCTCCCCAATTTTGCCGAGTGCTGGTGGGATCAAGTCATTCTGGACATCC
	TGTTGTGCAATGGCGGTGGCATTTGGCTGGGCATGGTCGTTTGCCGGTTTTTAGAGATGAGGACTTA
	CCACTGGGCAAGCTTCAAGGACATTCATACCACCACCGGGAAGATCAAGAGAGCTGTTCTGCAGTTC
	ACTCCTGCTAGCTGGACCTATGTTCGATGGTTTGACCCCAAATCTTCTTTTCAGAGAGTAGCTGGAG
	TGTACCTTTTCATGATCATCTGGCAGCTGACTGAGTTGAATACCTTCTTCTTGAAGCATATCTTTGT
	GTTCCAAGCCAGTCATCCATTAAGTTGGGGTAGAATTCTCTTTATTGGTGGCATCACAGCTCCCACA
	GTGAGACAGTACTACGCTTACCTCACCGACACACAGTGCAAGCGCGTAGGAACACAATGCTGGGTGT
	TTGGGGCTTTCACCACTTTCCTCTGTCTGTACGGCCATGATTTGGTATGCAGAACACTATGGTCACCG
	AGAAAAGACCTACTCGGAGTGTGAAGATGGCACCTACAGTCCAGAGATCTCCTGGCATCACAGGAAA
	GGGACAAAAGGTTCTGAAGACAGCCCACCCAAGCATGCAGGCAACAACGAAAGCCATTCTTCCAGGA
	GAAGGAATCGGCATTCCAAGTCAAAAGTCACCAATGGCGTTGGAAAGAAATGAAAAACCCTGGTTAA
	TCAAAGATGTTCCAGAGTGCCTAGCACTTGAGAGGGGAAATGGAACTCATTTGGAACTCCCCGTGAGGA
	GGTCGAGGCGCACAGGGCAAGCAGGAAGAGGCGAGGGCACTTGGGGGTCATTATTTGAGATCGTAAG
	TCTTGTTTCCCACAGACCTGGCCGCGTCAGGCAGATCATCGCCTGGGGTGGGCCTTTGCCAACGTGGGG
	TCTCTTCTAACTTCAGCACTTGACATGCGGTCACCGGTGGCAGCGCGGTGTGTTGAAGGGAAACGGT
	AGCTATTCATTCACAGTTGCCAAGAGCAGCTCCGCGCCTGCTGGATCGTGGATGCAGCGTAAACATC
	TTCCTTCAGACGAGGCATTAACCCCATGGTTAATGGACTGGTCACCAGTTTTTATTTTATTTTTATG
	AATCTACCTTTCCATTGATTGATTTAAGTTCAGGCCACTTTTCTGTCTTTTATTTGGTTACTGTTGT
	TATTTGTTTTTAAGTTAGGATGCTTTTTAACAGCCTTTAGAAGCCGCTGCTGAAATTGATACTGGGG
	GAAGGGTTCCCCTTCCTTCTAGAGCAGAAAAGGGAGAGAAGTGTTGTATTCCTGTTTGGTAACCTCA
	GTCTCCTGTAAGACCTCCTACCACATGGCGAGTATACACCAATCAGGAGAGGGTAGCTGCCTGCATA
	GGAGCCTCGCTTCCGATTATTCCCTTCCCAATATTATTCATCCAGACTTAGCCACAGTGCACAAAAG
	CAAACCTGCTAGAGAGGCAGTGAACACCACAGCTTCTCCCCAGCTTGGTGCCTTTTACATCGGGTTT
	GTTCTCCTTCCATGGTGTGTTGCTGACATTGTCACTGAGTCCCATGTGAGGTGCTGGTGAGTATTAC
	CTTTCATCTGTGCCATGCTCTAGAACCTTGACCTTGATAGTTCACCACGTCTGATGGATCCCTGTTT
	TAAATAAAAACGATTCACTTTAAAGCCT

	ORF Start: ATG at 4		ORF Stop: TGA at 1324

	SEQ ID NO: 182	440 aa	MW at 51772.5kD
NOV43a, CG152256-01 Protein Sequence	MASCVGSRTLSKDDVNYKMHFRMINEQQVEDITIDFFYRPHTITLLSFTIVSLMYFAFTRDDSVPED NIWRGILSVIFFFLIISVLAFPNGPFTRPHPALWRMVFGLSVLYFLFLVFLLPLNFEQVKSLMYWLD PNLRYATREADVMEYAVNCHVITWERIISHFDIFAFGHFWGWAMKALLIRSYGLCWTISITWELTEL FFMHLLPNFAECWWDQVILDILLCNGGGIWLGMVVCRFLEMRTYHWASFKDIHTTTGKIKRAVLQFT PASWTYVRWFDPKSSFQRVAGVYLFMIIWQLTELNTFFLKHIFVFQASHPLSWGRILFIGGITAPTV RQYYAYLTDTQCKRVGTQCWVFGAFTTFLCLYGMIWYAEHYGHREKTYSECEDGTYSPEISWHHRKG TKGSEDSPPKHAGNNESHSSRRRNRHSKSKVTNGVGKK		

Further analysis of the NOV43a protein yielded the following properties shown in Table 43B.

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Table 43B. Protein Sequence Properties NOV43a	
PSort analysis:	0.6000 probability located in plasma membrane; 0.4000 probability located in Golgi body; 0.3000 probability located in endoplasmic reticulum (membrane); 0.0300 probability located in mitochondrial inner membrane
SignalP analysis:	No Known Signal Sequence Predicted

A search of the NOV43a protein against the Geneseq database, a proprietary database that contains sequences published in patents and patent publication, yielded several homologous proteins shown in Table 43C.

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Table 43C. Geneseq Results for NOV43a				
Geneseq Identifier	Protein/Organism/Length [Patent #, Date]	NOV43a Residues/ Match Residues	Identities/ Similarities for the Matched Region	Expect Value
ABB89640	Human polypeptide SEQ ID NO 2016 - Homo sapiens, 473 aa. [WO200190304-A2, 29-NOV-2001]	1..440 1..473	440/473 (93%) 440/473 (93%)	0.0
AAB58945	Breast and ovarian cancer associated antigen protein sequence SEQ ID 653 - Homo sapiens, 516 aa. [WO200055173-A1, 21-SEP-2000]	1..440 44..516	439/473 (92%) 439/473 (92%)	0.0
ABB71324	Drosophila melanogaster polypeptide SEQ ID NO 40764 - Drosophila melanogaster, 498 aa. [WO200171042-A2, 27-SEP-2001]	3..359 59..412	206/357 (57%) 276/357 (76%)	e-133
AAB73515	Human transferase HTFS-22, SEQ ID NO:22 - Homo sapiens, 487 aa. [WO200132888-A2, 10-MAY-2001]	22..361 45..389	128/351 (36%) 185/351 (52%)	2e-60
AAM79907	Human protein SEQ ID NO 3553 - Homo sapiens, 529 aa. [WO200157190-A2, 09-AUG-2001]	22..361 63..407	128/351 (36%) 185/351 (52%)	2e-60

In a BLAST search of public sequence datbases, the NOV43a protein was found to have homology to the proteins shown in the BLASTP data in Table 43D.

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Table 43D. Public BLASTP Results for NOV43a

Protein Accession Number	Protein/Organism/Length	NOV43a Residues/ Match Residues	Identities/ Similarities for the Matched Portion	Expect Value
P48651	Phosphatidylserine synthase I (Serine-exchange enzyme I) (EC 2.7.8.-) - Homo sapiens (Human), 473 aa.	1..440 1..473	440/473 (93%) 440/473 (93%)	0.0
Q99LH2	Similar to phosphatidylserine synthase 1 - Mus musculus (Mouse), 473 aa.	1..440 1..473	428/473 (90%) 437/473 (91%)	0.0
Q00576	Phosphatidylserine synthase I (Serine-exchange enzyme I) (EC 2.7.8.-) - Cricetulus longicaudatus (Long-tailed hamster) (Chinese hamster), 471 aa.	1..440 1..471	428/473 (90%) 434/473 (91%)	0.0
O55024	Phosphatidylserine synthase-1 - Mus musculus (Mouse), 473 aa.	1..440 1..473	421/473 (89%) 432/473 (91%)	0.0
Q9BSY0	Similar to phosphatidylserine synthase 1 - Homo sapiens (Human), 334 aa (fragment).	145..440 6..334	292/329 (88%) 293/329 (88%)	e-178

PFam analysis predicts that the NOV43a protein contains the domains shown in the Table 43E.

Table 43E. Domain Analysis of NOV43a			
Pfam Domain	NOV43a Match Region	Identities/ Similarities for the Matched Region	Expect Value
COLFI	119..137	10/19 (53%) 14/19 (74%)	0.12
PSS	96..370	179/310 (58%) 267/310 (86%)	1.1e-206

Example 44.

The NOV44 clone was analyzed, and the nucleotide and encoded polypeptide sequences are shown in Table 44A.

Table 44A. NOV44 Sequence Analysis			
	SEQ ID NO: 183	1151 bp	
NOV44a, CG171804-01 DNA Sequence	CNTGNATTTGGCCGGGGGGGCCATGTAGCTCCGAGCGGCGGATCGCGAGCCTCCTGCGAACCCCAGCC TGCACGCCCGGTTAGCATTCGGCCGGGAGATGCGGCAGTGGAATCTGGAAGGGCGGTGAAAAACCTA CGTCCTGCCCTCGCCCGGCCTCTCCATTCGTCCCCCGGGTAGAGAGGGTCGGCTCGTGCTCATCATC CTGTGCTCCGTGGTCTTCTCTGCCGTCTACATCCTCCTGTGCTGCTGGGCCGGCCTGCCCCTCTGCC TGGCCACCTGCCTGGACCACCACTTCCCCACAGGCTCCAGGCCCACTGTGCCGGGACCCCTGCACTT CAGTGGATATAGCAGTGTGCCAGATGGGAAGCCGCTGGTCCGCGAGCCCTGCCGCAGCTGTGCCGTG GTGTCCAGCTCCGGCCAAATGCTGGGCTCAGGCCTGGGTGCTGAGATCGACAGTGCCGAGTGCGTGT TCCGCATGAACCAGGCGCCCACCGTGGGCTTTGAGGCGGATGTGGGCCAGCGCAGCACCCTGCGTGT CGTCTCACACACAAGCGTGCCGCTGCTGCTGCGCAACTATTCACACTACTTCCAGAAGGCCCGAGAC ACGCTCTACATGGTGTGGGGCCAGGGCAGGCACATGGACCGGGTGCTCGGCGGCCGCACCTACCGCA CGCTGCTGCAGCTCACCAGGATGTACCCCGGCCTGCAGGTGTACACCTTCACGGAGCGCATGATGGC CTACTGCGACCAGATCTTCCAGGACGGACGGGCAAGAACCGGAGGCAGTCGGGCTCCTTCCTCAGC ACCGGCTGGTTCACCATGATCCTCGCGCTGGAGCTGTGTGAGGAGATCGTGGTCTATGGGATGGTCA GCGACAGCTACTGCAGGGAGAAGAGCCACCCCTCAGTGCCTTACCACTACTTTGAGAAGGGCCGGCT AGATGAGTGTCAGATGTACCTGGCACACGAGCAGGCGCCCCGAAGCGCCCACCGCTTCATCACTGAG AAGGCGGTCTTCTCCCGCTGGGCCAAGAAGAGGCCCATCGTGTTCGCCCATCCGTCCTGGAGGACTG AGTAGCTTCCGTCGTCCTGCCAGCCGCCATGCCGTTGCGAGGCCTCCGGGATGTCCCATCCCAAGCC ATCACACTCCAC		
	ORF Start: ATG at 421		ORF Stop: TAG at 1075

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	SEQ ID NO: 184	218 aa	MW at 25333.8kD
NOV44a, CG171804-01 Protein Sequence	MLGSGLGAEIDSAECVFRMNQAPTVGFEADVGQRSTLRVVSHTSVPLLLRNYSHYFQKARDTLYMVW GQGRHMDRVLGGRTYRTLLQLTRMYPGLQVYTFTERMMAYCDQIFQDETGKNRRQSGSFLSTGWFTM ILALELCEEIVVYGMVSDSYCREKSHPSVPYHYFEKGRLDECQMYLAHEQAPRSAHRFITEKAVFSR WAKKRPIVFAHPSWRTE		

10   Further analysis of the NOV44a protein yielded the following properties shown in Table 44B.

Table 44B. Protein Sequence Properties NOV44a	
PSort analysis:	0.6400 probability located in microbody (peroxisome); 0.3000 probability located in nucleus; 0.2068 probability located in lysosome (lumen); 0.1000 probability located in mitochondrial matrix space
SignalP analysis:	No Known Signal Sequence Predicted

15

A search of the NOV44a protein against the Geneseq database, a proprietary database that contains sequences published in patents and patent publication, yielded several homologous proteins shown in Table 44C.

Table 44C. Geneseq Results for NOV44a				
Geneseq Identifier	Protein/Organism/Length [Patent #, Date]	NOV44a Residues/ Match Residues	Identities/ Similarities for the Matched Region	Expect Value
AAB75350	Human secreted protein #9 - Homo sapiens, 302 aa. [WO200100806-A2, 04-JAN-2001]	1..218 85..302	218/218 (100%) 218/218 (100%)	e-128
AAB61614	Human protein HP03380 - Homo sapiens, 302 aa. [WO200102563-A2, 11-JAN-2001]	1..218 85..302	218/218 (100%) 218/218 (100%)	e-128
AAB25764	Human secreted protein SEQ ID #76 - Homo sapiens, 302 aa. [WO200037491-A2, 29-JUN-2000]	1..218 85..302	218/218 (100%) 218/218 (100%)	e-128
AAB28674	Human carbohydrate-modifying enzyme Incyte ID No: 983984CD1 - Homo sapiens, 302 aa. [WO200063351-A2, 26-OCT-2000]	1..218 85..302	218/218 (100%) 218/218 (100%)	e-128
AAB24495	Human secreted protein sequence encoded by gene 5 SEQ ID NO:120 - Homo sapiens, 345 aa. [WO200035937-A1, 22-JUN-2000]	1..218 128..345	217/218 (99%) 217/218 (99%)	e-128

5   In a BLAST search of public sequence datbases, the NOV44a protein was found to have homology to the proteins shown in the BLASTP data in Table 44D.

Table 44D. Public BLASTP Results for NOV44a				
Protein Accession Number	Protein/Organism/Length	NOV44a Residues/ Match Residues	Identities/ Similarities for the Matched Portion	Expect Value

Q9H4F1	Alpha-N-acetyl-neuraminyl-2,3-beta-gala ctosyl-1,3-N- acetylgalactosaminide alpha-2,6-sialyltransferase (EC 2.4.99.7) (NeuAc-alpha-2,3-Gal-beta-1,3-GalNAc-a lpha-2, 6-sialyltransferase) (ST6GalNAc IV) (Sialyltransferase 7D) - Homo sapiens (Human), 302 aa.	1..218 85..302	218/218 (100%) 218/218 (100%)	e-128
Q9H4F1	Alpha2,6-sialyltransferase - Homo sapiens (Human), 302 aa.	1..218 85..302	217/218 (99%) 218/218 (99%)	e-128
Q9NWU6	CDNA FLJ20593 fis, clone KAT08984 - Homo sapiens (Human), 302 aa.	1..218 85..302	217/218 (99%) 217/218 (99%)	e-127
Q9UKU1	NeuAc-alpha-2,3-Gal-beta-1,3-GalNAc-al pha-2, 6-sialyltransferase alpha2,6-sialyltransferase - Homo sapiens (Human), 302 aa.	1..218 85..302	216/218 (99%) 216/218 (99%)	e-127
Q9R2B6	Alpha-N-acetyl-neuraminyl-2,3-beta-gala ctosyl-1,3-N- acetylgalactosaminide alpha-2,6-sialyltransferase (EC 2.4.99.7) (NeuAc-alpha-2,3-Gal-beta-1,3-GalNAc-a lpha-2, 6-sialyltransferase) (ST6GalNAc IV) (Sialyltransferase 7D) - Mus musculus (Mouse), 360 aa.	1..218 143..360	202/218 (92%) 207/218 (94%)	e-118

PFam analysis predicts that the NOV44a protein contains the domains shown in the Table 44E.

5

Table 44E. Domain Analysis of NOV44a			
Pfam Domain	NOV44a Match Region	Identities/ Similarities for the Matched Region	Expect Value
Glyco_transf_29	1..202	65/324 (20%) 184/324 (57%)	6e-43

Example 45.

10   The NOV45 clone was analyzed, and the nucleotide and encoded polypeptide sequences are shown in Table 45A.

Table 45A. NOV45 Sequence Analysis

	SEQ ID NO: 185	1475 bp	
NOV45a, CG171841-01 DNA Sequence	AGGACTCCAAGCGCCATGGCCGCTGCCGCCCGAGCCCGGGTCGCGTACTTGCTGAGGCAACTGCAAC GCGCAGCATGGCTGTTTCAAATATTAGATATGGAGCAGCAGTTACAAAGGAAGTAGGAATGGCAGAC CTAAAAAACATGGGTGCTAAAAATGTGTGCTTGATGACAGACAAGAACCTCTCCAAGCTCCCTCCTG TGCAAGTAGCTATGGATTCCCTAGTGAAGAATGGCATCCCCTTTACGGTTTATGATAATGTGAGAGT GGAACCAACGGATAGCTTCATGGAAGCTATTGAGTTTGCCCAAAAGGGAGCTTTTGATGCCTATGTT GCTGTCGGTGGTGGCTCTACCATGGACACCTGTAAGGCTGCTAATCTGTATGCATCCAGCCCTCATT CTGATTTCCTAGATTATGTCAGTGCCCCCATTGGCAAGGGAAAGCCTGTGTCTGTGCCTCTTAAGCC TCTGATTGCAGTGCCAACTACCTCAGGAACCGGGAGTGAAACTACTGGGGTTGCCATTTTTGACTAT GAACACTTGAAAGTAAAAATTGGCATCACTTCGAGAGCCATCAAACCCACACTGGGACTGATTGATC CTCTGCACACCCTCCACATGCCTGCCCGAGTGGTCGCCAACAGTGGCTTTGATGTGTTTAGCCATGC CCTGGAGTCATACACCACCCTGCCCTACCACCTGCGGAGCCCCTGCCCTTCAAATCCCATCACACGG CCTGCGTACCAGGGCAGCAACCCAATCAGTGACATTTGGGCTATCCACGCGCTGCGGATCGTGGCTA AGTATCTGAAGGCTGTCAGAAATCCCGATGATCTTGAAGCAAGGTCTCATATGCACTTGGCAAGTGC TTTTGCTGGCATCGGCTTTGGAAATGCTGGTGTTCATCTGCATGGAATGTCTTACCCAATTTCAGGT TTAGTGAAGATGTATAAAGCAAAGGATTACAATGTGGATCACCCACTGGTGCCCCATGGCCTTTCTG TGGTGCTCACGTCCCCAGCGGTGTTCACTTTCACGGCCCAGATGTTTCCAGAGCGACACCTGGAGAT GGCAGAACTTCTAGGAGCCGACACCCGCACTGCCAGGATCCAAGATGCAGGGCTGGTGTTGGCAGAC ACGCTCCGGAAATTCTTATTCGATCTGGATGTTGATGATGGCCTAGCAGCTGTTGGTTACTCCAAAG CTGATATCCCCGCACTAGTGAAAGGAACGCTGCCCCAGGAAAGGGTCACCAAGCTTGCACCCTGTCC CCAGTCAGAAGAGGATCTGGCTGCTCTGTTTGAAGCTTCAATGAAACTGTATTAATTGTCATTTTAA CTGAAAGAATTACCGCTGGCCATTGTAGTGCTGAGAGCAAGAGCTGATCTAGCTAGGGCTTTGTCTT TTCATCTTTGCGCATAACTTACCTGTTACCAGTATAGGTGGGATATACATTTATCTTGCAGGAAATT C
	ORF Start: ATG at 75		ORF Stop: TAA at 1326

	SEQ ID NO: 186	417 aa	MW at 44871.2kD
NOV45a, CG171841-01 Protein Sequence	MAVSNIRYGAAVTKEVGMADLKNMGAKNVCLMTDKNLSKLPPVQVAMDSLVKNGIPFTVYDNVRVEP TDSFMEAIEFAQKGAFDAYVAVGGGSTMDTCKAANLYASSPHSDFLDYVSAPIGKGKPVSVPLKPLI AVPTTSGTGSETTGVAIFDYEHLKVKIGITSRAIKPTLGLIDPLHTLHMPARVVANSGFDVFSHALE SYTTLPYHLRSPCPSNPITRPAYQGSNPISDIWAIHALRIVAKYLKAVRNPDDLEARSHMHLASAFA GIGFGNAGVHLHGMSYPISGLVKMYKAKDYNVDHPLVPHGLSVVLTSPAVFTFTAQMFPERHLEMAE LLGADTRTARIQDAGLVLADTLRKFLFDLDVDDGLAAVGYSKADIPALVKGTLPQERVTKLAPCPQS EEDLAALFEASMKLY

Further analysis of the NOV45a protein yielded the following properties shown in Table 45B.

Table 45B. Protein Sequence Properties NOV45a	
PSort analysis:	0.4500 probability located in cytoplasm; 0.3188 probability located in microbody (peroxisome); 0.2355 probability located in lysosome (lumen); 0.1000 probability located in mitochondrial matrix space
SignalP analysis:	No Known Signal Sequence Predicted

A search of the NOV45a protein against the Geneseq database, a proprietary database that contains sequences published in patents and patent publication, yielded several homologous proteins shown in Table 45C.

Table 45C. Geneseq Results for NOV45a				
Geneseq Identifier	Protein/Organism/Length [Patent #, Date]	NOV45a Residues/ Match Residues	Identities/ Similarities for the Matched Region	Expect Value
AAE21522	Human dehydrogenase DHDR-6 protein - Homo sapiens, 467 aa. [WO200216562-A2, 28-FEB-2002]	1..417 49..467	413/420 (98%) 414/420 (98%)	0.0
AAB73686	Human oxidoreductase protein ORP-19 - Homo sapiens, 467 aa. [WO200144448-A2, 21-JUN-2001]	1..417 49..467	412/420 (98%) 413/420 (98%)	0.0
ABB59876	Drosophila melanogaster polypeptide SEQ ID NO 6420 - Drosophila melanogaster, 464 aa. [WO200171042-A2, 27-SEP-2001]	1..417 46..464	254/420 (60%) 327/420 (77%)	e-146
ABG08093	Novel human diagnostic protein #8084 - Homo sapiens, 268 aa. [WO200175067-A2, 11-OCT-2001]	62..322 1..268	240/268 (89%) 243/268 (90%)	e-131
AAB42855	Human ORFX ORF2619 polypeptide sequence SEQ ID NO:5238 - Homo sapiens, 212 aa. [WO200058473-A2, 05-OCT-2000]	247..417 41..212	168/172 (97%) 170/172 (98%)	7e-91

5   In a BLAST search of public sequence datbases, the NOV45a protein was found to have homology to the proteins shown in the BLASTP data in Table 45D.

Table 45D. Public BLASTP Results for NOV45a				
Protein Accession Number	Protein/Organism/Length	NOV45a Residues/ Match Residues	Identities/ Similarities for the Matched Portion	Expect Value

CAD28993	Sequence 4 from Patent WO0216562 - Homo sapiens (Human), 467 aa.	1..417 49..467	413/420 (98%) 414/420 (98%)	0.0
Q96MF9	CDNA FLJ32430 fis, clone SKMUS2001129, weakly similar to NAD-dependent methanol dehydrogenase (EC 1.1.1.244) - Homo sapiens (Human), 419 aa.	1..417 1..419	412/420 (98%) 413/420 (98%)	0.0
Q8R0N6	Hypothetical 45.0 kDa protein - Mus musculus (Mouse), 419 aa.	1..417 1..419	372/420 (88%) 394/420 (93%)	0.0
Q9W265	T3DH protein - Drosophila melanogaster (Fruit fly), 464 aa.	1..417 46..464	254/420 (60%) 327/420 (77%)	e-145
Q95S86	GM05887p - Drosophila melanogaster (Fruit fly), 425 aa.	1..417 7..425	254/420 (60%) 327/420 (77%)	e-145

PFam analysis predicts that the NOV45a protein contains the domains shown in the Table 45E.

Table 45E. Domain Analysis of NOV45a			
Pfam Domain	NOV45a Match Region	Identities/ Similarities for the Matched Region	Expect Value
Fe-ADH	4..205	68/216 (31%) 143/216 (66%)	5.6e-28
Fe-ADH	228..288	30/68 (44%) 51/68 (75%)	2.5e-10

Example 46.

The NOV46 clone was analyzed, and the nucleotide and encoded polypeptide sequences are shown in Table 46A.

Table 46A. NOV46 Sequence Analysis			
	SEQ ID NO: 187	1310 bp	
NOV46a, CG173017-01 DNA Sequence	CTACTCTCAGCCAGGAATCATGTCTTGGGCCGCTCGCCCGCCCTTCCTCCCTCAGCGGCATGCCGCA GGGCAGTGTGGGCCGGTGGGGGTGCGAAAAGAAATGCATTGTGGGGTCGCGTCCCGGTGGCGGCGGC GACGGCCCTGGCTGGATCCCGCAGCGGCGGCGGCGGCGGCGGTGGCAGGCGGAGAACAACAAACCCC GGAGCCGGAGCCAGGGGAGGCTGGACGGGACGGGGATGGGCGACAGCGGGCGGGGTGGCCCTGGGGCT GGCAAACGGCTATGTGCAATCTGCCGGGGACAGAAGCTCAGGCAAACACTACGGGGTTTACAGCTGTG AGGGTTGCAAGGGCTTCTTCAAACGCACCATCCGCAAAGACCTTACATACTCTTGCCGGGACAACAA AGACTGCACAGTGGACAAGCGCCAGCGGAACCGCTGTCAGTACTGCCGCTATCAGAAGTGCCTGGCC ACTGGCATGAAGAGGGGAGGCGGTACAGGAGGAGCGTCAGCGGGGAAAGGACAGGGATGGGGATGGGG AGGGGGCTGGGGGGAGCCCCCGAGGAGATGCCTGTGGACAGGATCCTGGAGGCAGAGCTTGCTGTGGA ACAGAAGAGTGACCAGGGCGTTGAGGGTCCTGGGGGAACCGGGGGTAGCGGCAGCAGCCCAAATGAC CCTGTGACTAACATCTGTCAGGCAGCTGACAAACAGCTATTCACGCTTGTTGAGTGGGCGAAGAGGA TCCCACACTTTTCCTCCTTGCCTCTGGATGATCAGGTCATATTGCTGCGGGCAGGCTGGAATGAACT CCTCATTGCCTCCTTTTCACACCGATCCATTGATGTTCAGGATGGCATCCTCCTTGCCACAGGTCTT CACGTGCACCGCCAACTCAGCCCATTCAGCAGGAGTAGGAGCCATCTTTGATCGGGTGCTGACAGAGC TAGTGTCCAAAATGCGTGACATGAGGATGGACAAGACAGAGCTTGGCTGCCTGAGGGCAATCATTCT GTTTAATCCAGATGCCAAGGGCCTCTCCAACCCTAGTGAGGTGGAGGTCCTGCGGGAGAAAGTGTAT GCATCACTGGAGACCTACTGCAAACAGAAGTACCCTGAGCAGCAGGGACGGTTTGCCAAGCTGCTGC TACGTCTTCCTGCCCTCCGGTCCATTGGCCTTAAGTGTCTAGAGCATCTGTTTTTCTTCAAGCTCAT TGGTGACACCCCCATCGACACCTTCCTCATGGAGATGCTTGAGGCTCCCCATCAACTGGCCTGAGCT CAGACCCAGACGTGGTGCTTCTCCACACTGGAGGAGC		
	ORF Start: ATG at 20		ORF Stop: TGA at 1268

	SEQ ID NO: 188	416 aa	MW at 45778.7kD
NOV46a, CG173017-01 Protein Sequence	MSWAARPPFLPQRHAAGQCGPVGVRKEMHCGVASRWRRRRPWLDPAAAAAAAVAGGEQQTPEPEPGE AGRDGMGDSGRGGPGAGKRLCAICGDRSSGKHYGVYSCEGCKGFFKRTIRKDLTYSCRDNKDCTVDK RQRNRCQYCRYQKCLATGMKREAVQEERQRGKDRDGDGEGAGGAPEEMPVDRILEAELAVEQKSDQG VEGPGGTGGSGSSPNDPVTNICQAADKQLFTLVEWAKRIPHFSSLPLDDQVILLRAGWNELLIASFS HRSIDVRDGILLATGLHVHRNSAHSAGVGAIFDRVLTELVSKMRDMRMDKTELGCLRAIILFNPDAK GLSNPSEVEVLREKVYASLETYCKQKYPEQQGRFAKLLLRLPALRSIGLKCLEHLFFFKLIGDTPID TFLMEMLEAPHQLA		

Further analysis of the NOV46a protein yielded the following properties shown in Table 46B.

Table 46B. Protein Sequence Properties NOV46a	
PSort analysis:	0.9700 probability located in nucleus; 0.3000 probability located in microbody (peroxisome); 0.1000 probability located in mitochondrial matrix space; 0.1000 probability located in lysosome (lumen)
SignalP analysis:	No Known Signal Sequence Predicted

A search of the NOV46a protein against the Geneseq database, a proprietary database that contains sequences published in patents and patent publication, yielded several homologous proteins shown in Table 46C.

Table 46C. Geneseq Results for NOV46a

Geneseq Identifier	Protein/Organism/Length [Patent #, Date]	NOV46a Residues/ Match Residues	Identities/ Similarities for the Matched Region	Expect Value
AAU78297	Human Retinoid X Receptor beta (RXRbeta) protein - Homo sapiens, 533 aa. [WO200218420-A2, 07-MAR-2002]	41..416 156..533	346/378 (91%) 352/378 (92%)	0.0
AAR72483	Human H-2RIIBP - Homo sapiens, 533 aa. [US5403925-A, 04-APR-1995]	41..416 156..533	346/378 (91%) 352/378 (92%)	0.0
AAR39468	hRXR-beta1 - Homo sapiens, 533 aa. [WO9315216-A, 05-AUG-1993]	41..416 156..533	346/378 (91%) 352/378 (92%)	0.0
AAR39469	hRXR-beta2 - Homo sapiens, 510 aa. [WO9315216-A, 05-AUG-1993]	41..416 133..510	345/378 (91%) 351/378 (92%)	0.0
AAY21625	Ligand binding domain of nuclear receptor hRXRbeta - Homo sapiens, 525 aa. [WO9926966-A2, 03-JUN-1999]	41..416 148..525	345/378 (91%) 351/378 (92%)	0.0

5 In a BLAST search of public sequence datbases, the NOV46a protein was found to have homology to the proteins shown in the BLASTP data in Table 46D.

Table 46D. Public BLASTP Results for NOV46a

Protein Accession Number	Protein/Organism/Length	NOV46a Residues/ Match Residues	Identities/ Similarities for the Matched Portion	Expect Value
S37781	retinoid X receptor beta - human, 533 aa.	41..416 156..533	346/378 (91%) 352/378 (92%)	0.0
Q95L53	Retinoid X receptor beta - Mustela vison (American mink), 525 aa (fragment).	41..416 148..525	346/378 (91%) 352/378 (92%)	0.0

P28702	Retinoic acid receptor RXR-beta - Homo sapiens (Human), 533 aa.	41..416 156..533	346/378 (91%) 352/378 (92%)	0.0
A41651	retinoic acid receptor coregulator - rat, 451 aa.	41..416 74..451	341/378 (90%) 349/378 (92%)	0.0
D41727	retinoid X receptor beta - mouse, 448 aa.	41..416 71..448	341/378 (90%) 349/378 (92%)	0.0

PFam analysis predicts that the NOV46a protein contains the domains shown in the Table 46E.

Table 46E. Domain Analysis of NOV46a			
Pfam Domain	NOV46a Match Region	Identities/ Similarities for the Matched Region	Expect Value
zf-C4	86..161	49/77 (64%) 73/77 (95%)	1.5e-54
hormone_rec	227..409	74/207 (36%) 157/207 (76%)	3.3e-68

Example 47.

The NOV47 clone was analyzed, and the nucleotide and encoded polypeptide sequences are shown in Table 47A.

Table 47A. NOV47 Sequence Analysis			
	SEQ ID NO: 189	1229 bp	
NOV47a, CG173347-01 DNA Sequence	CCGAGACCATGGGGAAGCTCGTGGCGCTGGTCCTGCTGGGGGTCGGCCTGTCCTTAGTCGGGGAGAT GTTTCTGGCGTTTAGAGAAAGGGTGAATGCCTCTCGAGAAGTGGAGCCAGTAGAACCTGAAAACTGC CACCTTATTGAGGAACTTGAAAGTGGCTCTGAAGATATTGATATACTTCCTAGTGGGCTGGCTTTTA TCTCCAGTCTGCAGGTCTGTTGGAGTTTGCTGGAAGTCCACTCCAGACCCTGTTTGCCTGGGTATCA CCAGTGGAGGCTGCAGAACGGCAAATATTGCTGCCTGATTTTTCTTCTGGAAGCTTCATCCCAGAGG GGCATCCGCCTGTATGAGGGATTAAAATATCCAGGCATGCCAAACTTTGCGCCAGATGAACCAGGAA AAATCTTCTTGATGGATCTGAATGAACAAAACCCAAGGGCACAAGCACTAGAAATCAGTGGTGGATT TGACAAAGAATTATTTAATCCACATGGGATCAGTATTTTCATCGACAAAGACAATACTGTGTATCTT TATGTTGTGAATCATCCCCACATGAAGTCCACTGTGGAGATATTTAAATTTGAGGAACAACAACGTT CTCTGGTATACCTGAAAACTATAAAACATGAACTTCTCAAAAGTGTGAATGACATTGTGGTTCTTGG ACCAGAACAGTTCTATGCCACCAGAGACCACTATTTTACCAACTCCCTCCTGTCATTTTTTGAGATG ATCTTGGATCTTCGCTGGACTTATGTTCTTTTCTACAGCCCAAGGGAGGTTAAAGTGGTGGCCAAAG GATTTTGTAGTGCCAATGGGATCACAGTCTCAGCAGACCAGAAGTATGTCTATGTAGCTGATGTAGC AGCTAAGAACATTCACATAATGGAAAAACATGATAACTGGGATTTAACTCAACTGAAGGTGATACAG TTGGGCACCTTAGTGGATAACCTGACTGTCGATCCTGCCACAGGAGACATTTTGGCAGGATGCCATC CTAATCCTATGAAGCTACTGAACTATAACCCTGAGGACCCTCCAGGATCAGAAGTACTTCGCATCCA GAATGTTTTGTCTGAGAAGCCCAGGGTGAGCACCGTGTATGCCAACAATGGCTCTGTGCTTCAGGGC ACCTCTGTGGCTTCTGTGTGTACCATGGGAAAATTCTCATAGGCACCGTATTTCNCAAAACTCTGTACT

	GTGAGCTC**TAGA**CTCT**AGATAGT**		
	ORF Start: ATG at 9		ORF Stop: TAG at 1215

	SEQ ID NO: 190	402 aa	MW at 45160.5kD
NOV47a, CG173347-01 Protein Sequence	MGKLVALVLLGVGLSLVGEMFLAFRERVNASREVEPVEPENCHLIEELESGSEDIDILPSGLAFISS LQVCWSLLEVHSRPCLPGYHQWRLQNGKYCCLIFLLEASSQRGIRLYEGLKYPGMPNFAPDEPGKIF LMDLNEQNPRAQALEISGGFDKELFNPHGISIFIDKDNTVYLYVVNHPHMKSTVEIFKFEEQQRSLV YLKTIKHELLKSVNDIVVLGPEQFYATRDHYFTNSLLSFFEMILDLRWTYVLFYSPREVKVVAKGFC SANGITVSADQKYVYVADVAAKNIHIMEKHDNWDLTQLKVIQLGTLVDNLTVDPATGDILAGCHPNP MKLLNYNPEDPPGSEVLRIQNVLSEKPRVSTVYANNGSVLQGTSVASVYHGKILIGTVFXKTLYCEL		

Further analysis of the NOV47a protein yielded the following properties shown in Table 47B.

10

Table 47B. Protein Sequence Properties NOV47a	
PSort analysis:	0.8200 probability located in outside; 0.1900 probability located in lysosome (lumen); 0.1000 probability located in endoplasmic reticulum (membrane); 0.1000 probability located in endoplasmic reticulum (lumen)
SignalP analysis:	Cleavage site between residues 31 and 32

A search of the NOV47a protein against the Geneseq database, a proprietary
15 database that contains sequences published in patents and patent publication, yielded several homologous proteins shown in Table 47C.

Table 47C. Geneseq Results for NOV47a				
Geneseq Identifier	Protein/Organism/Length [Patent #, Date]	NOV47a Residues/ Match Residues	Identities/ Similarities for the Matched Region	Expect Value
ABB97287	Novel human protein SEQ ID NO: 555 - Homo sapiens, 354 aa. [WO200222660-A2, 21-MAR-2002]	1..402 1..354	352/402 (87%) 352/402 (87%)	0.0
AAG75494	Human colon cancer antigen protein SEQ ID NO:6258 - Homo sapiens, 370 aa. [WO200122920-A2, 05-APR-2001]	2..402 18..370	352/401 (87%) 352/401 (87%)	0.0

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ABG08350	Novel human diagnostic protein #8341 - Homo sapiens, 382 aa. [WO200175067-A2, 11-OCT-2001]	1..402 24..382	330/407 (81%) 333/407 (81%)	e-178
AAU11925	Protein sequence of rabbit paraoxonase-3 (PON3) mutant D324N - Oryctolagus cuniculus, 355 aa. [WO200190336-A2, 29-NOV-2001]	1..402 1..355	294/403 (72%) 318/403 (77%)	e-164
AAU11922	Protein sequence of rabbit paraoxonase-3 (PON3) mutant N169D - Oryctolagus cuniculus, 355 aa. [WO200190336-A2, 29-NOV-2001]	1..402 1..355	294/403 (72%) 318/403 (77%)	e-164

In a BLAST search of public sequence datbases, the NOV47a protein was found to have homology to the proteins shown in the BLASTP data in Table 47D.

5

Table 47D. Public BLASTP Results for NOV47a				
Protein Accession Number	Protein/Organism/Length	NOV47a Residues/ Match Residues	Identities/ Similarities for the Matched Portion	Expect Value
Q15166	Serum paraoxonase/arylesterase 3 (EC 3.1.1.2) (EC 3.1.8.1) (PON 3) (Serum aryldiakylphosphatase 3) (A-esterase 3) (Aromatic esterase 3) - Homo sapiens (Human), 354 aa.	1..402 1..354	354/402 (88%) 354/402 (88%)	0.0
Q9BZH9	Paraoxanase-3 - Homo sapiens (Human), 354 aa (fragment).	1..402 1..354	351/402 (87%) 351/402 (87%)	0.0
Q9BGN0	Paraoxonase 3 - Oryctolagus cuniculus (Rabbit), 354 aa.	1..402 1..354	293/402 (72%) 318/402 (78%)	e-164

| Q62087 | Serum paraoxonase/arylesterase 3 (EC 3.1.1.2) (EC 3.1.8.1) (PON 3) (Serum aryldiakylphosphatase 3) (A-esterase 3) (Aromatic esterase 3) - Mus musculus (Mouse), 354 aa. | 1..402 1..354 | 283/402 (70%) 314/402 (77%) | e-158 |
| Q90952 | Serum paraoxonase/arylesterase 2 (EC 3.1.1.2) (EC 3.1.8.1) (PON 2) (Serum aryldiakylphosphatase 2) (A-esterase 2) (Aromatic esterase 2) - Gallus gallus (Chicken), 354 aa. | 1..402 1..354 | 230/402 (57%) 287/402 (71%) | e-131 |

PFam analysis predicts that the NOV47a protein contains the domains shown in the Table 47E.

Table 47E. Domain Analysis of NOV47a			
Pfam Domain	NOV47a Match Region	Identities/ Similarities for the Matched Region	Expect Value
Arylesterase	2..402	230/422 (55%) 348/422 (82%)	1.2e-190

Example 48.

The NOV48 clone was analyzed, and the nucleotide and encoded polypeptide sequences are shown in Table 48A.

Table 48A. NOV48 Sequence Analysis			
	SEQ ID NO: 191	2109 bp	
NOV48a, CG56234-01 DNA Sequence	CCTTCCATACCTCCCCGGCTCCGCTCGGTTCCTGGCCACCCCGCAGCCCCTGCCCAGGTGCCATGGC CGCATTGTACCGCCCTGGCCTGCGGCTTAACTGGCATGGGCTGAGCCCCTTGGGCTGGCCATCATGC CGTAGCATCCAGACCCTGCGAGTGCTTAGTGGAGATCTGGGCCAGCTTCCCACTGGCATTCGAGATT TTGTAGAGCACAGTGCCCGCCTGTGCCAACCAGAGGGCATCCACATCTGTGATGGAACTGAGGCTGA GAATACTGCCACACCTGACCCTGCTGGAGCAGCAGGGCCTCATCCGAAAGCTCCCCAAGTACAATAAC TGCTGGCTGGCCCCGCACAGACCCCAAGGATGTGGCACGAGTAGAGAGCAAGACGGTGATTGTAACTC CTTCTCAGCGGGACACGGTACCACTCCCGCCTGGTGGGGCCCGTGGGCAGCTGGGCAACTGGATGTC CCCAGCTGATTTCCAGCGAGCTGTGGATGAGAGGTTTCCAGGCTGCATGCAGGGCCGCACCATGTAT GTGCTTCCATTCAGCATGGGTCCTGTGGGCTCCCCGCTGTCCCGCATCGGGGTGCAGCTCACTGACT CAGCCTATGTGGTGGCAAGCATGCGTATTATGACCCGACTGGGGACACCTGTGCTTCAGGCCCTGGG AGATGGTGACTTTGTCAAGTGTCTGCACTCCGTGGGCCAGCCCCTGACAGGACAAGGGGAGCCAGTG

279

	AGCCAGTGGCCGTGCAACCCAGAGAAAACCCTGATTGGCCACGTGGCCGACCAGCGGGAGATCATCT CCTTCGGCAGCGGCTATGGTGGCAACTCCCTGCTGGGCAAGAAGTGCTTTGCCCTACGCATCGCCTC TCGGCTGGCCCGGGATGAGGGCTGGCTGGCAGAGCACATGCTGATCCTGGGCATCACCAGCCCTGCA GGGAAGAAGCGCTATGTGGCAGCCGCCTTCCCTAGTGCCTGTGGCAAGACCAACCTGGCTATGATGC GGCCTGCACTGCCAGGCTGGAAAGTGGAGTGTGTGGGGGATGATATTGCTTGGATGAGGTTTGACAG TGAAGGTCGACTCCGGGCCATCAACCCTGAGAACGGCTTCTTTGGGGGTTGCCCCTGGTACCTCTGCC ACCACCAATCCCAACGCCATGGCTACAATCCAGAGTAACACTATTTTTACCAATGTGGCTGAGACCA GTGATGGTGGCGTGTACTGGGAGGGCATTGACCAGCCTCTTCCACCTGGTGTTACTGTGACCTCCTG GCTGGGCAAACCCTGGAAATCTGGTGACAAGGAGCCCTGTGCACATCCCAACTCTCGATTTTGTGCC CCGGCTCGCCAGTGCCCCATCATGGACCCAGCCTGGGAGGCCCCAGAGGGTGTCCCCATTGACGCCA TCATCTTTGGTGGCCGCAGACCCAAAGGGGTACCCCTGGTATACGAAGCCTTCAACTGGCGTCATGG GGTGTTTGTGGGCAGCGCCATGCGCTCTGAGTCCACTGCTGCAGCAGAACACAAAGGGAAGATCATC ATGCACGACCCATTTGCCATGCGGCCCTTTTTTGGCTACAACTTCGGGCACTACCTGGAACACTGGC TGAGCATGGAAGGGCGCAAGGGGGCCCAGCTGCCCCGTATCTTCCATGTCAACTGGTTCCGGCGTGA CGAGGCAGGGCACTTCCTGTGGCCAGGCTTTGGGGAGAATGCTCGGGTGCTGGATCTGCCCGG CGGTTAGAGGGGGAGGACAGTGCCCGAGAGACACCCATTGGGCTGGTACCAAAGGAAGGAGCCTTGG ATCTCAGCGGCCTCAGAGCTATAGACACCACTCAGCTGTTCTCCCTCCCCAAGGACTTCTGGGAACA GGAGGTTCGTGACATTCGGAGCTACCTGACAGAGCAGGTCAACCAGGATCTGCCCAAAGAGGTGTTG GCTGAGCTTGAGGCCCTGGAGAGACGTGTGCACAAAATGTGA<u>CCTGAGGCCCTAGTCTAGCAAGAGG ACATAGCACCCTCATCTGGGAATAGGGAAGGCACCCTTGCAGAAAATATGAGCAATTTGATATTAACT AACATCTTCAATGTGCCATAGACCTTCCCACA</u>
	ORF Start: ATG at 63		ORF Stop: TGA at 1983

	SEQ ID NO: 192	640 aa	MW at 70688.2kD
NOV48a, CG56234-01 Protein Sequence	MAALYRPGLRLNWHGLSPLGWPSCRSIQTLRVLSGDLGQLPTGIRDFVEHSARLCQPEGIHICDGTE AENTATLTLLEQQGLIRKLPKYNNCWLARTDPKDVARVESKTVIVTPSQRDTVPLPPGGARGQLGNW MSPADFQRAVDERFPGCMQGRTMYVLPFSMGPVGSPLSRIGVQLTDSAYVVASMRIMTRLGTPVLQA LGDGDFVKCLHSVGQPLTGQGEPVSQWPCNPEKTLIGHVPDQREIISFGSGYGGNSLLGKKCFALRI ASRLARDEGWLAEHMLILGITSPAGKKRYVAAAFPSACGKTNLAMMRPALPGWKVECVGDDIAWMRF DSEGRLRAINPENGFFGVAPGTSATTNPNAMATIQSNTIFTNVAETSDGGVYWEGIDQPLPPGVTVT SWLGKPWKSGDKEPCAHPNSRFCAPARQCPIMDPAWEAPEGVPIDAIIFGGRRPKGVPLVYEAFNWR HGVFVGSAMRSESTAAAEHKGKIIMHDPFAMRPFGYNFGHYLEHWLSMEGRKGAQLPRIFHVNWFR RDEAGHFLWPGFGENARVLDWICRRLEGEDSARETPIGLVPKEGALDLSGLRAIDTTQLFSLPKDFW EQEVRDIRSYLTEQVNQDLPKEVLAELEALERRVHKM

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	SEQ ID NO: 193	2069 bp	
NOV48b, CG56234-02 DNA Sequence	CCCGCCTTCCATACCTCCCCGGCTCCGCTCGGTTCCTGGCCACCCCGCAGCCCCTGCCCAGGTGCCA TGGCCGCATTGTACCGCCCTGGCCTGCGGCTTAACTGGCATGGGCTGAGCCCCTTGGGCTGGCCATC ATGCCGTAGCATCCAGACCCTGCGAGTGCTTAGTGGAGATCTGGGCCAGCTTCCCACTGGCATTCGA GATTTTGTAGAGCACAGTGCCCGCCTGTGCCAACCAGAGGGCATCCACATCTGTGATGGAACTGAGG CTGAGAATACTGCCACACTGACCCTGCTGGAGCAGCAGGGCCTCATCCGAAAGCTCCCCAAGTACAA TAACTGCTGGCTGGCCCGCACAGACCCCAAGGATGTGGCACGAGTAGAGAGCAAGACGGTGATTGTA ACTCCTTCTCAGCGGGACACGGTACCACTCCCGCCTGGTGGGGCCTGTGGGCAGCTGGGCAACTGGA TGTCCCCAGCTGATTTCCAGCGAGCTGTGGATGAGAGGTTTCCAGGCTGCATGCAGGGCCGCACCAT GTATGTGCTTCCATTCAGCATGGGTCCTGTGGGCTCCCCGCTGTCCCGCATCGGGGTGCAGCTCACT GACTCAGCCTATGTGGTGGCAAGCATGCGTATTATGACCCGACTGGGGACCCTGTGCTTCAGGCCC TGGGAGATGGTGACTTTGTCAAGTGTCTGCACTCCGTGGGCCAGCCCCTGACAGGACAAGGGGAGCC AGTGAGCCAGTGGCCGTGCAACCCAGAGAAAACCCTGATTGGCCACGTGCCCGACCAGCGGGAGATC ATCTCCTTCGGCAGCGGCTATGGTGGCAACTCCCTGCTGGGCAAGAAGTGCTTTGCCCTACGCATCG CCTCTCGGCTGGCCCGGGATGAGGGCTGGCTGGCAGAGCACATGCTGATCCTGGGCATCACCAGCTC TGCAGGGAAGAAGCGCTATGTGGCAGCCGCCTTCCCTAGTGCCTGTGGCAAGACCAACCTGGCTATG ATGCGGCCTGCACTGCCAGGCTGGAAAGTGGAGTGTGTGGGGGATGATATTGCTTGGATGAGGTTTG ACAGTGAAGGTCGACTCCGGGCCATCAACCCTGAGAACGGCTTCTTTGGGGGTTGCCCCTGGTACCTC TGCCACCACCAATCCCAACGCCATGGCTACAATCCAGAGTAACACTATTTTTACCAATGTGGCTGAG ACCAGTGATGGTGGCGTGTACTGGGAGGGCATTGACCAGCCTCTTCCACCTGGTGTTACTGTGACCT CCTGGCTGGGCAAACCCTGGAAACCTGGTGACAAGGAGCCCTGTGCACATCCCAACTCTCGATTTTG TGCCCCGGCTCGCCAGTGCCCCATCATGGACCCAGCCTGGGAGGCCCCAGAGGGTGTCCCCATTGAC GCCATCATCTTTGGTGGCCGCAGACCCAAAGGGGAAGATCATCATGCACGACCCATTTGCCATGCGGC CCTTTTTTGGCTACAACTTCGGGCACTACCTGGAACACTGGCTGAGCATGGAAGGGCGCAAGGGGGC CCAGCTGCCCCGTATCTTCCATGTCAACTGGTTCCGGCGTGACGAGGCAGGGCACTTCCTGTGGCCA GGCTTTGGGGAGAATGCTCGGGTGCTAGACTGGATCTGCCGGCGGTTAGAGGGGGAGGACAGTGCCC

	GAGAGACACCCATTGGGCTGGTGCCAAAGGAAGGAGCCTTGGATCTGAGCGGCGTCAGAGGTATAGA CACCACTCAGCTGTTCTCCCTCCCCAAGGACTTCTGGGAACAGGAGGTTCGTGACATTCGGAGCTAC CTGACAGAGCAGGTCAACCAGGATCTGCCCAAAGAGGTGTTGGCTGAGCTTGAGGCCCTGGAGAGAC GTGTGCACAAAATGTGACCTGAGGCCTAGTCTAGCAAGAGGACATAGCACCCTCATCTGGGAATAGG GAAGGCACCTTGCAGAAAATATGAGCAATTGATATTAACTAACATCTTCAATGTGCCATAGACCTTC CCACAAAGACTGTCCAATAATAAGAGATGCTTATCTATTTTAAAAAAAAAAAAAAAAAAA
ORF Start: ATG at 67		ORF Stop: TGA at 1891

	SEQ ID NO: 194	608 aa	MW at 67027.1kD
NOV48b, CG56234-02 Protein Sequence	MAALYRPGLRLNWHGLSPLGWPSCRSIQTLRVLSGDLGQLPTGIRDFVEHSARLCQPEGIHICDGTE AENTATLTLLEQQGLIRKLPKYNNCWLARTDPKDVARVESKTVIVTPSQRDTVPLPPGGACGQLGNW MSPADFQRAVDERFPGCMQGRTMYVLPFSMGPVGSPLSRIGVQLTDSAYVVASMRIMTRLGTPVLQA LGDGDFVKCLHSVGQPLTGQGEPVSQWPCNPEKTLIGHVPDQREIISFGSGYGGNSLLGKKCFALRI ASRLARDEGWLAEHMLILGITSPAGKKALCAAAFPSACGKTNLAMMRPALPGWKVECVGDDIAWMRF DSEGRLRAINPENGFFGVAPGTSATTNPNAMATIQSNTIFTNVAETSDGGVYWEGIDQPLPPGVTVT SWLGKPWKPGDKEPCAHPNSRFCAPARQCPIMDPAWEAPEGVPIDAIIFGGRRPKGKIIMHDPFAMR PFFGYNFGHYLEHWLSMEGRKGAQLPRIFHVNWFRRDEAGHFLWPGFGENARVLDWICRRLEGEDSA RETPIGLVPKEGALDLSGLRAIDTTQLFSLPKDFWEQEVRDIRSYLTEQVNQDLPKEVLAELEALER RVHKM

Sequence comparison of the above protein sequences yields the following sequence relationships shown in Table 48B.

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Table 48B. Comparison of NOV48a against NOV48b.		
Protein Sequence	**NOV48a Residues/ Match Residues**	**Identities/ Similarities for the Matched Region**
NOV48b	1..640 1..608	577/640 (90%) 577/640 (90%)

Further analysis of the NOV48a protein yielded the following properties shown in

15   Table 48C.

Table 48C. Protein Sequence Properties NOV48a	
PSort analysis:	0.6402 probability located in microbody (peroxisome); 0.3000 probability located in nucleus; 0.2412 probability located in lysosome (lumen); 0.1000 probability located in mitochondrial matrix space
SignalP analysis:	No Known Signal Sequence Predicted

A search of the NOV48a protein against the Geneseq database, a proprietary
database that contains sequences published in patents and patent publication, yielded
several homologous proteins shown in Table 48D.

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Table 48D. Geneseq Results for NOV48a				
Geneseq Identifier	Protein/Organism/Length [Patent #, Date]	NOV48a Residues/ Match Residues	Identities/ Similarities for the Matched Region	Expect Value
AAY80296	Human mitochondrial phosphoenolpyruvate carboxykinase SEQ ID NO:1 - Homo sapiens, 640 aa. [US6030837-A, 29-FEB-2000]	1..640 1..640	634/640 (99%) 634/640 (99%)	0.0
AAB71890	Mouse PEPCK-cytosolic protein - Mus musculus, 622 aa. [US6187545-B1, 13-FEB-2001]	31..640 14..622	440/610 (72%) 519/610 (84%)	0.0
AAB71880	Human PEPCK-cytosolic protein - Homo sapiens, 622 aa. [US6187545-B1, 13-FEB-2001]	31..640 14..622	438/610 (71%) 517/610 (83%)	0.0
ABB65318	Drosophila melanogaster polypeptide SEQ ID NO 22746 - Drosophila melanogaster, 647 aa. [WO200171042-A2, 27-SEP-2001]	27..640 35..647	394/616 (63%) 480/616 (76%)	0.0
ABB65322	Drosophila melanogaster polypeptide SEQ ID NO 22758 - Drosophila melanogaster, 638 aa. [WO200171042-A2, 27-SEP-2001]	30..640 29..638	402/613 (65%) 469/613 (75%)	0.0

In a BLAST search of public sequence datbases, the NOV48a protein was found to
have homology to the proteins shown in the BLASTP data in Table 48E.

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Table 48E. Public BLASTP Results for NOV48a

Protein Accession Number	Protein/Organism/Length	NOV48a Residues/ Match Residues	Identities/ Similarities for the Matched Portion	Expect Value
Q16822	Phosphoenolpyruvate carboxykinase, mitochondrial precursor [GTP] (EC 4.1.1.32) (Phosphoenolpyruvate carboxylase) (PEPCK-M) - Homo sapiens (Human), 640 aa.	1..640 1..640	635/640 (99%) 635/640 (99%)	0.0
S69546	phosphoenolpyruvate carboxykinase (GTP) (EC 4.1.1.32) precursor, mitochondrial - human, 640 aa.	1..640 1..640	634/640 (99%) 634/640 (99%)	0.0
Q91Z10	Similar to phosphoenolpyruvate carboxykinase 2 (mitochondrial) - Mus musculus (Mouse), 640 aa.	1..640 1..640	590/640 (92%) 609/640 (94%)	0.0
Q8R3X7	Similar to RIKEN cDNA 9130022B02 gene - Mus musculus (Mouse), 535 aa (fragment).	106..640 1..535	504/535 (94%) 518/535 (96%)	0.0
P07379	Phosphoenolpyruvate carboxykinase, cytosolic [GTP] (EC 4.1.1.32) (Phosphoenolpyruvate carboxylase) (PEPCK-C) - Rattus norvegicus (Rat), 622 aa.	31..640 14..622	441/610 (72%) 520/610 (84%)	0.0

PFam analysis predicts that the NOV48a protein contains the domains shown in the Table 48F.

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Table 48F. Domain Analysis of NOV48a			
Pfam Domain	NOV48a Match Region	Identities/ Similarities for the Matched Region	Expect Value
PEPCK	46..640	445/608 (73%) 591/608 (97%)	0

Example 49.

The NOV49 clone was analyzed, and the nucleotide and encoded polypeptide
sequences are shown in Table 49A.

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Table 49A. NOV49 Sequence Analysis			
	SEQ ID NO: 195	1202 bp	
NOV49a, CG56836-01 DNA Sequence	TGTAAGCGATCTGGTTCCCACCTCAGCCTCCCGAGTAGTGTCTTCAGGCCTATGGAGAGCAGCTTGC GTGGGCTGGGCCTGCAGTACCTGGTTTGCATAGATGATTGGCAGGTGGATCTAGGATCCGGCTTCCA ACATGTGGCAGCTCTGGGCCTCCCTCTGCTGCCTGCTGGTGTTGGCCAATGCCCGGAGCAGGCCCTC TTTCCATCCCCTGTCGGATGAGCTGGTCAACTATGTCAACAAACGGAATACCACGTGGCAGGCCGGG CACAACTTCTACAACGTGGACATGAGCTACTTGAAGAGGCTATGTGGTACCTTCCTGGGTGGGCCCA AGCCACCCCAGAGAGTTATGTTTACCGAGGACCTGAAGCTGCCTGCAAGCTTCGATGCACGGGAACA ATGGCCACAGTGTCCCACCATCAAAGAGATCAGAGACCAGGGCTCCTGTGGCTCCTGCTGGGCCTTC GGGGCTGTGGAAGCCATCTCTGACCGGATCTGCATCCACACCAATGCGCACGTCAGCGTGGAGGTGT CGGCGGAGGACCTGCTCACATGCTGTGGCAGCATGTGTGGGGACGGCTGTAATGGTGGCTATCCTGC TGAAGCTTGGAACTTCTGGACAAGAAAGGCCTGGTTTCTGGTGGCCTCTATGAATCCCATGTAGGG TGCAGACCGTACTCCATCCCTCCCTGTGAGCACCACGTCAACGGCTCCCGGCCCCCATGCACGGGGG AGGGAGATACCCCCAAGTGTAGCAAGATCTGTGAGCCTGGCTACAGCCCGACCTACAAACAGGACAA GCACTACGGATACAATTCCTACAGCGTCTCCAATAGCGAGAAGGACATCATGGCCGAGATCTACAAA AACGGCCCCGTGGAGGGAGCTTTCTCTGTGTATTCGGACTTCCTGCTCTACAAGTCAGGAGTGTACC AACACGTCACCGGAGAGATGATGGGTGGCCATGCCATCCGCATCCTGGGCTGGGGAGTGGAGAATGG CACACCCTACTGGCTGGTTGCCAACTCCTGGAACACTGACTGGGGTGACAATGGCTTCTTTAAAATA CTCAGAGGACAGGATCACTGTGGAATCGAATCAGAAGTGGTGGCTGGAATTCCACGCACCGATCAGT ACTGGGAAAAGATCTAATCTGCCGTGGGCCTGTCGTGCCAGTCCTGGGGGCGAGATCGGGGTA
	ORF Start: ATG at 137		ORF Stop: TAA at 1154

	SEQ ID NO: 196	339 aa	MW at 37821.3kD
NOV49a, CG56836-01 Protein Sequence	MWQLWASLCCLLVLANARSRPSFHPLSDELVNYVNKRNTTWQAGHNFYNVDMSYLKRLCGTFLGGPK PPQRVMFTEDLKLPASFDAREQWPQCPTIKEIRDQGSCGSCWAFGAVEAISDRICIHTNAHVSVEVS AEDLLTCCGSMCGDGCNGGYPAEAWNFWTRKGLVSGGLYESHVGCRPYSIPPCEHHVNGSRPPCTGE GDTPKCSKICEPGYSPTYKQDKHYGYNSYSVSNSEKDIMAEIYKNGPVEGAFSVYSDFLLYKSGVYQ HVTGEMMGGHAIRILGWGVENGTPYWLVANSWNTDWGDNGFFKILRGQDHCGIESEVVAGIPRTDQY WEKI

	SEQ ID NO: 197	723 bp	
NOV49b, CG56836-02 DNA Sequence	ACATGGTGGATCTAGGATCCGGCTTCCAACATGTGGCAGCTCTGGGCCTCCCTCTGCTGCCTGCTGG TGTTGGCCAATGCCCGGAGCAGGCCCTCTTTTCCATCCCCTGTCGGATGAGCTGGTCAACTATGTCAA CAAACGGAATACCACGTGGCAGGCCGGGCACAACTTCTACAACGTGGACATGAGCTACTTGAAGAGG CTATGTGGTACCTTCCTGGGTGGGCCCAAGCCACCCCAGAGAGTTATGTTTACCGAGGACCTGAAGC TGCCTGCAAGCTTCGATGCACGGGAACAATGGCCACAGTGTCCCACCATCAAAGAGATCAGAGACCA GGGCTCCTGTGGCTCCTGCTGGGCCTTCGGGGGCTGTGGAAGCCATCTCTGACCGGATCTGCATCCAC ACCAATGCGCACGTCAGCGTGGAGGTGTCGGCGGAGGACCTGCTCACCTGCTCACGTGCTTCTACAAGTCAG GAGTGTACCAACACGTCACCGGAGAGATGATGGGTGGCCATGCCATCCGCATCCTGGGCTGGGGAGT GGAGAATGGCACACCCTACTGGCTGGTTGCCAACTCCTGGAACACTGACTGGGGTGACAATGGCTTC TTTAAAATACTCAGAGGACAGGATCACTGTGGAATCGAATCAGAAGTGGTGGCTGGAATTCCACGCA CCGATCAGTACTGGGAAAAGATCTAATCTGCCGTGGGCCTGTCGTGCCAAACC
	ORF Start: ATG at 31		ORF Stop: TAA at 694

	SEQ ID NO: 198	221 aa	MW at 24974.2kD
NOV49b, CG56836-02 Protein Sequence	MWQLWASLCCLLVLANARSRPSFHPLSDELVNYVNKRNTTWQAGHNFYNVDMSYLKRLCGTFLGGPK PPQRVMFTEDLKLPASFDAREQWPQCPTIKEIRDQGSCGSCWAFGAVEAISDRICIHTNAHVSVEVS AEDLLTCLLYKSGVYQHVTGEMMGGHAIRILGWGVENGTPYWLVANSWNTDWGDNGFFKILRGQDHC GIESEVVAGIPRTDQYWEKI		

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	SEQ ID NO: 199	1028 bp	
NOV49c, CG56836-03 DNA Sequence	TGTAAGCGATCTGGTTCCCACCTCAGCCTCCCGAGTAGTGTCTTCAGGCCTATGGAGAGCAGCTTGC GTGGGCTGGGCCTGCAGTACCTGGTTTGCATAGATGATTGGCAGGTGGATCTAGGATCCGGCTTCCA ACATGTGGCAGCTCTGGGCCTCCCTCTGCTGCCTGCTGGTGTTGGCCAATGCCCGGAGCAGGCCCTC TTTCCATCCCCTGTCGGATGAGCTGGTCAACTATGTCAACAAACGGAATACCACGTGGCAGGCCGGG CACAACTTCTACAACGTGGACATGAGCTACTTGAAGAGGCTATGTGGTACCTTCCTGGGTGGGCCCA AGCCACCCCAGAGAGTTATGTTTACCGAGGACCTGAAGCTGCCTGCAAGCTTCGATGCACGGGAACA ATGGCCACAGTGTCCCACCATCAAAGAGATCAGAGACCAGGGCTCCTGTGGCTCCTGCTGGGCCTTC CGGCGGAGGACCTGCTCACATGCTGTGGCAGCATGTGTGGGGACGGCTGTAATGGTGGCTATCCTGC TGAAGCTTGGAACTTCTGGACAAGAAAAGGCCTGGTTTCTGGTGGCCTCTATGAATCCAATAGCGAG AAGGACATCATGGCCGAGATCTACAAAAACGGCCCCGTGGAGGGAGCTTTCTCTGTGTATTCGGACT TCCTGCTCTACAAGTCAGGAGTGTACCAACACGTCACCGGAGAGATGATGGGTGGCCATGCCATCCG CATCCTGGGCTGGGGAGTGGAGAATGGCACACCCTACTGGCTGGTTGCCAACTCCTGGAACACTGAC TGGGGTGACAATGGCTTCTTTAAAATACTCAGAGGACAGGATCACTGTGGAATCGAATCAGAAGTGG TGGCTGGAATTCCACGCACCGATCAGTACTGGGAAAAGATCTAATCTGCCGTGGGCCTGTCGTGCCA GTCCTGGGGGCGAGATCGGGGTA		
	ORF Start: ATG at 137		ORF Stop: TAA at 980

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	SEQ ID NO: 200	281 aa	MW at 31423.2kD
NOV49c, CG56836-03 Protein Sequence	MWQLWASLCCLLVLANARSRPSFHPLSDELVNYVNKRNTTWQAGHNFYNVDMSYLKRLCGTFLGGPK PPQRVMFTEDLKLPASFDAREQWPQCPTIKEIRDQGSCGSCWAFGAVEAISDRICIHTNAHVSVEVS AEDLLTCCGSMCGDGCNGGYPAEAWNFWTRKGLVSGGLYESNSEKDIMAEIYKNGPVEGAFSVYSDF LLYKSGVYQHVTGEMMGGHAIRILGWGVENGTPYWLVANSWNTDWGDNGFFKILRGQDHCGIESEVV AGIPRTDQYWEKI		

15

	SEQ ID NO: 201	1028 bp	
NOV49d, CG56836-04 DNA Sequence	TGTAAGCGATCTGGTTCCCACCTCAGCCTCCCGAGTAGTGTCTTCAGGCCTATGGAGAGCAGCTTGC GTGGGCTGGGCCTGCAGTACCTGGTTTGCATAGATGATTGGCAGGTGGATCTAGGATCCGGCTTCCA ACATGTGGCAGCTCTGGGCCTCCCTCTGCTGCCTGCTGGTGTTGGCCAATGCCCGGAGCAGGCCCTC TTTCCATCCCCTGTCGGATGAGCTGGTCAACTATGTCAACAGACGGAATACCACGTGGCAGGCCGGG CACAACTTCTACAACGTGGACATGAGCTACTTGAAGAGGCTATGTGGTACCTTCCTGGGTGGGCCCA AGCCACCCCAGAGAGTTATGTTTACCGAGGACCTGAAGCTGCCTGCAAGCTTCGATGCACGGGAACA ATGGCCACAGTGTCCCACCATCAAAGAGATCAGAGACCAGGGCTCCTGTGGCTCCTGCTGGGTTTCT GGTGGCCTCTATGAATCCCATGTAGGGTGCAGACCGTACTCCATCCCTCCCTGTGAGCACCACGTCA ACGGCTCCCGGCCCCCATGCACGGGGGAGGGAGATACCCCCAAGTGTAGCAAGATCTGTGAGCCTGG CTACAGCCCGACCTACAAACAGGACAAGCACTACGGATCAATTCCTACAGCGTCTCCAATAGCGAG AAGGACATCATGGCCGAGATCTACAAAAACGGCCCCGTGGAGGGAGCTTTCTCTGTGTATTCGGACT TCCTGCTCTACAAGTCAGGAGTGTACCAACACGTCACCGGAGAGATGATGGGTGGCCATGCCATCCG CATCCTGGGCTGGGGAGTGGAGAATGGCACACCCTACTGGCTGGTTGCCAACTCCTGGAACACTGAC TGGGGTGACAATGGCTTCTTTAAAATACTCAGAGGACAGGATCACTGTGGAATCGAATCAGAAGTGG TGGCTGGAATTCCACGCACCGATCAGTACTGGGAAAAGATCTAATCTGCCGTGGGCCTGTCGTGCCA GTCCTGGGGGCGAGATCGGGGTA		

	ORF Start: ATG at 137		ORF Stop: TAA at 980

	SEQ ID NO: 202	281 aa	MW at 31732.5kD
NOV49d, CG56836-04 Protein Sequence	MWQLWASLCCLLVLANARSRPSFHPLSDELVNYVNRRNTTWQAGHNFYNVDMSYLKRLCGTFLGGPK PPQRVMFTEDLKLPASFDAREQWPQCPTIKEIRDQGSCGSCWVSGGLYESHVGCRPYSIPPCEHHVN GSRPPCTGEGDTPKCSKICEPGYSPTYKQDKHYGYNSYSVSNSEKDIMAEIYKNGPVEGAFSVYSDF LLYKSGVYQHVTGEMMGGHAIRILGWGVENGTPYWLVANSWNTDWGDNGFFKILRGQDHCGIESEVV AGIPRTDQYWEKI		

	SEQ ID NO: 203	340 bp	
NOV49e, 247856403 DNA Sequence	AGGCTCCGCGGCCGCCCCCTTCACCGGATCCCTGCCTGCAAGCTTCGATGCACGGGAACAATGGCCA CAGTGTCCCACCATCAAAGAGATCAGAGACCAGGGCTCCTGTGGCTCCTGCTGGGCCTTCGGGGCTG TGGAAGCCATCTCTGACCGGATCTGCATCCACACCAATGCGCACGTCAGCGTGGAGGTGTCGGCGGA GGACCTGCTCACATGCTGTGGCAGCATGTGTGGGGACGGCTGTAATGGTGGCTATCCTGCTGAAGCT TGGAACTTCTGGACAAGAAAAGGCCTGGTTTCTGGTGGCCTCTATCTCGAGGGCAAGGGTGGGCGCG CCGAC		
	ORF Start: at 2		ORF Stop: end of sequence

	SEQ ID NO: 204	113 aa	MW at 11834.0kD
NOV49e, 247856403 Protein Sequence	GSAAAPFTGSLPASFDAREQWPQCPTIKEIRDQGSCGSCWAFGAVEAISDRICIHTNAHVSVEVSAE DLLTCCGSMCGDGCNGGYPAEAWNFWTRKGLVSGGLYLEGKGGRAD		

	SEQ ID NO: 205	376 bp	
NOV49f, 247856434 DNA Sequence	AGGCTCCGCGGCCGCCCCCTTCACCGGATCCTCCAATAGCGAGAAGGACATCATGGCCGAGATCTAC AAAAACGGCCCCGTGGAGGGAGCTTTCTCTGTGTATTCGGACTTCCTGCTCTACAAGTCAGGAGTGT ACCAACACGTCACCGGAGAGATGATGGGTGGCCATGCCATCCGCATCCTGGGCTGGGGAGTGGAGAA TGGCACACCCTACTGGCTGGTTGCCAACTCCTGGAACACTGACTGGGGTGACAATGGCTTCTTTAAA ATACTCAGAGGACAGGATCACTGTGGAATCGAATCAGAAGTGGTGGCTGGAATTCCACGCACCGATC AGTACTGGGAAAAGATCCTCGAGGGCAAGGGTGGGCGCC		
	ORF Start: at 2		ORF Stop: end of sequence

	SEQ ID NO: 206	125 aa	MW at 13666.1kD
NOV49f, 247856434 Protein Sequence	GSAAAPFTGSSNSEKDIMAEIYKNGPVEGAFSVYSDFLLYKSGVYQHVTGEMMGGHAIRILGWGVEN GTPYWLVANSWNTDWGDNGFFKILRGQDHCGIESEVVAGIPRTDQYWEKILEGKGGRA		

	SEQ ID NO: 207	574 bp	
NOV49g, 247856497 DNA Sequence	AGGCTCCGCGGCCGCCCCCTTCACCGGATCCATGTGGCAGCTCTGGGCCTCCCTCTGCTGCCTGCTG GTGTTGGCCAATGCCCGGAGCAGGCCCTCTTTCCATCCCCTGTCGGATGAGCTGGTCAACTATGTCA ACAAACGGAATACCACGTGGCAGGCCGGGCACAACTTCTACAACGTGGACATGAGCTACTTGAAGAG GCTATGTGGTACCTTCCTGGGTGGGCCCAAGCCACCCCAGAGAGTTATGTTTACCGAGGACCTGAAG CTGCCTGCAAGCTTCGATGCACGGGAACAATGGCCACAGTGTCCCACCATCAAAGAGATCAGAGACC AGGGCTCCTGTGGCTCCTGCTGGGCCTTCGGGGCTGTGGAAGCCATCTCTGACCGGATCTGCATCCA CACCAATGCGCACGTCAGCGTGGAGGTGTCGGCGGAGGACCTGCTCACATGCTGTGGCAGCATGTGT GGGGACGGCTGTAATGGTGGCTATCCTGCTGAAGCTTGGAACTTCTGGACAAGAAAAGGCCTGGTTT CTGGTGGCCTCTATCTCGAGGGCAAGGGTGGGCGCGCC		
	ORF Start: at 2		ORF Stop: end of sequence

	SEQ ID NO: 208	191 aa	MW at 20877.5kD
NOV49g, 247856497 Protein Sequence	GSAAAPFTGSMWQLWASLCCLLVLANARSRPSFHPLSDELVNYVNKRNTTWQAGHNFYNVDMSYLKR LCGTFLGGPKPPQRVMFTEDLKLPASFDAREQWPQCPTIKEIRDQGSCGSCWAFGAVEAISDRICIH TNAHVSVEVSAEDLLTCCGSMCGDGCNGGYPAEAWNFWTRKGLVSGGLYLEGKGGRA		

	SEQ ID NO: 209	590 bp	
NOV49h, 247856493 DNA Sequence	AGGCTCCGCGGCCGCCCCCTTCACCGGATCCATGTGGCAGCTCTGGGCCTCCCTCTGCTGCCTGCTG GTGTTGGCCAATGCCCGGAGCAGGCCCTCTTTCCATCCCGTGTCGGATGAGCTGGTCAACTATGTCA ACAAACGGAATACCACGTGGCAGGCCGGGCACAACTTCTACAACGTGGACATGGGCTACTTGAAGAG CTGCCTGCAAGCTTCGATGCACGGGAACAATGGCCACAGTGTCCCACCATCAAAGAGATCAGAGACC AGGGCTCCTGTGGCTCCTGCTGGGCCTTCGGGGCTGTGGAAGCCATCTCTGACCGGATCTGCATCCA CACCAATGCGCACGTCAGCGTGGAGGTGTCGGCGGAGGACCTGCTCACCTGCTGTGGCAGCATGTGT CTGGTGGCCTCTATCTCGAGGGCAAGGGTGGGCGCGCCGACCCAGCTTTCTGTA		
	ORF Start: at 2		ORF Stop: end of sequence

	SEQ ID NO: 210	197 aa	MW at 21367.0kD
NOV49h, 247856493 Protein Sequence	GSAAAPFTGSMWQLWASLCCLLVLANARSRPSFHPVSDELVNYVNKRNTTWQAGHNFYNVDMGYLKR LCGTFLGGPKPPQRVMFTEDLKLPASFDAREQWPQCPTIKEIRDQGSCGSCWAFGAVEAISDRICIH TNAHVSVEVSAEDLLTCCGSMCGDGCNGGYPAEAWNFWTRKGLVSGGLYLEGKGGRADPAFCX		

	SEQ ID NO: 211	551 bp	
NOV49i,	AGGCTCCGCGGCCGCCCCCTTCACCGGATCCCGGAGCAGGCCCTCTTTCCATCCCCTGTCGGATGAG		

287

247856574 DNA Sequence	CTGGTCAACTATGTCAACAAACGGAATACCACGTGGCAGGCCGGGCACAACTTCTACAACGTGGACA TGAGCTACTTGAAGAGGCTATGTGGTACCTTCCTGGGTGGGCCCAAGCCACCCCAGAGAGTTATGTT TACCGAGGACCTGAAGCTGCCTGCAAGCTTCGATGCACGGGAACAATGGCCACAGTGTCCCACCATC AAAGAGATCAGAGACCAGGGCTCCTGTGGCTCCTGCTGGGCCTTCGGGGCTGTGGAAGCCATCTCTG ACCGGATCTGCATCCACACCAATGCGCACGTCAGCGTGGAGGTGTCGGCGGAGGACCTGCTCACATG CTGTGGCAGCATGTGTGGGGACGGCTGTAATGGTGGCTATCCTGCTGAAGCTTGGAACTTCTGGACA AGAAAAGGCCTGGTTTCTGGTGGCCTCTATCTCGAGGGCAAGGGTGGGCGCCCCGACCCAGCTTTCC CGTACAAAGCTGGCA
	ORF Start: at 2 ||ORF Stop: end of sequence

	SEQ ID NO: 212	184 aa	MW at 19933.2kD
NOV49i, 247856574 Protein Sequence	GSAAAPFTGSRSRPSFHPLSDELVNYVNKRNTTWQAGHNFYNVDMSYLKRLCGTFLGGPKPPQRVMF TEDLKLPASFDAREQWPQCPTIKEIRDQGSCGSCWAFGAVEAISDRICIHTNAHVSVEVSAEDLLTC CGSMCGDGCNGGYPAEAWNFWTRKGLVSGGLYLEGKGGRPDPAFPYKAGX		

	SEQ ID NO: 213	523 bp
NOV49j, 247856545 DNA Sequence	AGGCTCCGCGGCCGCCCCCTTCACCGGATCCCGGAGCAGGCCCTCTTTCCATCCCCTGTCGGATGAG CTGGTCAACTATGTCAACAAACGGAATACCACGTGGCAGGCCGGGCACAACTTCTACAACGTGGACA TGAGCTACTTGAAGAGGCTATGTGGTACCTTCCTGGGTGGGCCCAAGCCACCCCTGAGAGTTATGTT TACCGAGGACCTGAAGCTGCCTGCAAGCTTCGATGCACGGGAACAATGGCCACAGTGTCCCACCATC AAAGAGATCAGAGACCAGGGCTCCTGTGGCTCCTGCTGGGCCTTCGGGGCTGTGGAAGCCATCTCTG ACCGGATCTGCATCCACACCAATGCGCACGTCAGCGTGGAGGTGTCGGCGGAGGACCTGCTCACATG CTGTGGCGGCATGTGTGGGGACGGCTGTAATGGTGGCTATCCTGCTGAAGCTTGGAACTTCTGGACA AGAAAAGGCCTGGTTTCTGGTGGCCTCTATCTCGAGGGCAAGGGTGGGCGCGCC	
	ORF Start: at 2 ||ORF Stop: end of sequence

	SEQ ID NO: 214	174 aa	MW at 18915.1kD
NOV49j, 247856545 Protein Sequence	GSAAAPFTGSRSRPSFHPLSDELVNYVNKRNTTWQAGHNFYNVDMSYLKRLCGTFLGGPKPPLRVMF TEDLKLPASFDAREQWPQCPTIKEIRDQGSCGSCWAFGAVEAISDRICIHTNAHVSVEVSAEDLLTC CGGMCGDGCNGGYPAEAWNFWTRKGLVSGGLYLEGKGGRA		

	SEQ ID NO: 215	1036 bp
NOV49k, 275480714 DNA Sequence	CACCCTCGAGATGTGGCAGCTCTGGGCCTCCCTCTGCTGCCTGCTGGTGTTGGCCAATGCCCGGAGC AGGCCCTCTTTCCATCCCCTGTCGGATGAGCTGGTCAACTATGTCAACAAACGGAATACCACGTGGC AGGCCGGGCACAACTTCTACAACGTGGACATGAGCTACTTGAAGAGGCTATGTGGTACCTTCCTGGG TGGGCCCAAGCCACCCCAGAGAGTTATGTTTACCGAGGACCTGAAGCTGCCTGCAAGCTTCGATGCA CGGGAACAATGGCCACAGTGTCCCACCATCAAAGAGATCAGAGACCAGGGCTCCTGTGGCTCCTGCT GGGCCTTCGGGGCTGTGGAAGCCATCTCTGACCGGATCTGCATCCACACCAATGCGCACGTCAGCGT GGAGGTGTCGGCGGAGGACCTGCTCACATGCTGTGGCAGCATGTGTGGGGACGGCTGTAATGGTGGC TATCCTGCTGAAGCTTGGAACTTCTGGACAAGAAAAGGCCTGGTTTCTGGTGGCCTCTATGAATCCC ATGTAGGGTGCAGACCGTACTCCATCCCTCCCTGTGAGCACCACGTCAACGGCTCCCGGCCCCCATG	

| | CACGGGGGAGGGAGATACCCCCAAGTGTAGCAAGATCTGTGAGCCTGGCTACAGCCCGACCTACAAA CAGGACAAGCACTACGGATACAATTCCTACAGCGTCTCCAATAGCGAGAAGGACATCATGGCCGAGA TCTACAAAAACGGCCCCGTGGAGGGAGCTTTCTCTGTGTATTCGGACTTCCTGCTCTACAAGTCAGG AGTGTACCAACACGTCACCGGAGAGATGATGGGTGGCCATGCCATCCGCATCCTGGGCTGGGGAGTG GAGAATGGCACACCCTACTGGCTGGTTGCCAACTCCTGGAACACTGACTGGGGTGACAATGGCTTCT TTAAAATACTCAGAGGACAGGATCACTGTGGAATCGAATCAGAAGTGGTGGCTGGAATTCCACGCAC CGATCAGTACTGGGAAAAGATCGTCGACGGC |
| | ORF Start: at 2 | ORF Stop: end of sequence |

	SEQ ID NO: 216 .	345 aa	MW at 38435.9kD
NOV49k, 275480714 Protein Sequence	TLEMWQLWASLCCLLVLANARSRPSFHPLSDELVNYVNKRNTTWQAGHNFYNVDMSYLKRLCGTFLG GPKPPQRVMFTEDLKLPASFDAREQWPQCPTIKEIRDQGSCGSCWAFGAVEAISDRICIHTNAHVSV EVSAEDLLTCCGSMCGDGCNGGYPAEAWNFWTRKGLVSGGLYESHVGCRPYSIPPCEHHVNGSRPPC TGEGDTPKCSKICEPGYSPTYKQDKHYGYNSYSVSNSEKDIMAEIYKNGPVEGAPSVYSDPLLYKSG VYQHVTGEMMGGHAIRILGWGVENGTPYWLVANSWNTDWGDNGFFKILRGQDHCGIESEVVAGIPRT DQYWEKIVDG		

Sequence comparison of the above protein sequences yields the following sequence relationships shown in Table 49B.

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Table 49B. Comparison of NOV49a against NOV49b through NOV49k.		
Protein Sequence	**NOV49a Residues/ Match Residues**	**Identities/ Similarities for the Matched Region**
NOV49b	1..141 1..141	141/141 (100%) 141/141 (100%)
NOV49c	1..176 1..176	175/176 (99%) 176/176 (99%)
NOV49d	1..339 1..281	279/339 (82%) 280/339 (82%)
NOV49e	80..180 11..111	96/101 (95%) 96/101 (95%)
NOV49f	233..339 11..117	107/107 (100%) 107/107 (100%)
NOV49g	1..180 11..190	175/180 (97%) 175/180 (97%)
NOV49h	1..180 11..190	173/180 (96%) 174/180 (96%)
NOV49i	17..181 10..174	159/165 (96%) 160/165 (96%)
NOV49j	17..180 10..173	144/164 (87%) 145/164 (87%)

| NOV49k | 1..339 | 339/339 (100%) |
| | 4..342 | 339/339 (100%) |

Further analysis of the NOV49a protein yielded the following properties shown in Table 49C.

Table 49C. Protein Sequence Properties NOV49a	
PSort analysis:	0.3700 probability located in outside; 0.1900 probability located in lysosome (lumen); 0.1376 probability located in microbody (peroxisome); 0.1000 probability located in endoplasmic reticulum (membrane)
SignalP analysis:	Cleavage site between residues 18 and 19

A search of the NOV49a protein against the Geneseq database, a proprietary database that contains sequences published in patents and patent publication, yielded several homologous proteins shown in Table 49D.

Table 49D. Geneseq Results for NOV49a				
Geneseq Identifier	**Protein/Organism/Length [Patent #, Date]**	**NOV49a Residues/ Match Residues**	**Identities/ Similarities for the Matched Region**	**Expect Value**
AAR90616	Anti-procathepsin B monoclonal antibody - Homo sapiens, 339 aa. [JP07309900-A, 28-NOV-1995]	1..339 1..339	338/339 (99%) 339/339 (99%)	0.0
AAB53470	Human colon cancer antigen protein sequence SEQ ID NO:1010 - Homo sapiens, 344 aa. [WO200055351-A1, 21-SEP-2000]	1..339 6..344	338/339 (99%) 338/339 (99%)	0.0
ABP41147	Human ovarian antigen HOFMP73, SEQ ID NO:2279 - Homo sapiens, 346 aa. [WO200200677-A1, 03-JAN-2002]	1..339 8..346	290/339 (85%) 317/339 (92%)	0.0
ABB06116	Human NS protein sequence SEQ ID NO:208 - Homo sapiens, 273 aa. [WO200206315-A2, 24-JAN-2002]	1..267 1..267	266/267 (99%) 266/267 (99%)	e-167
ABB65378	Drosophila melanogaster polypeptide SEQ ID NO 22926 - Drosophila melanogaster, 340 aa. [WO200171042-A2, 27-SEP-2001]	13..331 13..339	190/330 (57%) 232/330 (69%)	e-113

In a BLAST search of public sequence datbases, the NOV49a protein was found to
have homology to the proteins shown in the BLASTP data in Table 49E.

Table 49E. Public BLASTP Results for NOV49a				
Protein Accession Number	**Protein/Organism/Length**	**NOV49a Residues/ Match Residues**	**Identities/ Similarities for the Matched Portion**	**Expect Value**

P07858	Cathepsin B precursor (EC 3.4.22.1) (Cathepsin B1) (APP secretase) - Homo sapiens (Human), 339 aa.	1..339 1..339	338/339 (99%) 339/339 (99%)	0.0
KHBOB	cathepsin B (EC 3.4.22.1) precursor - bovine, 335 aa.	1..335 1..335	280/335 (83%) 307/335 (91%)	e-180
P07688	Cathepsin B precursor (EC 3.4.22.1) - Bos taurus (Bovine), 335 aa.	1..335 1..335	279/335 (83%) 307/335 (91%)	e-180
P00787	Cathepsin B precursor (EC 3.4.22.1) (Cathepsin B1) (RSG-2) - Rattus norvegicus (Rat), 339 aa.	1..336 1..336	265/336 (78%) 299/336 (88%)	e-175
P10605	Cathepsin B precursor (EC 3.4.22.1) (Cathepsin B1) - Mus musculus (Mouse), 339 aa.	1..336 1..336	267/336 (79%) 297/336 (87%)	e-174

PFam analysis predicts that the NOV49a protein contains the domains shown in the Table 49F.

Table 49F. Domain Analysis of NOV49a			
Pfam Domain	**NOV49a Match Region**	**Identities/ Similarities for the Matched Region**	**Expect Value**
Peptidase_C1	80..329	112/344 (33%) 218/344 (63%)	1.3e-117

Example 50.

The NOV50 clone was analyzed, and the nucleotide and encoded polypeptide sequences are shown in Table 50A.

Table 50A. NOV50 Sequence Analysis			
	SEQ ID NO: 217	960 bp	
NOV50a, CG57284-01 DNA Sequence	CCCGTCCGAGCCCCGGCCCCAAGTAACGCCGCCGCCCCGGAGCCGCCTTGGAGGTCCCCCTCCCCAC TAAGTGCCTCTTTGCATAGCACCAGTCCCCACCCGCACGCTCTCTGGACCACTACAGCTGGACGGGC AATGGCGGGTCGGGGAGGCGCACGACGACCCAATGGACCAGCTGCTGGGAACAAGATCTGTCAATTT AAGCTGGTTCTGCTGGGGGAGTCTGCGGTAGGCAAATCCAGCCTCGTCCTCCGCTTTGTCAAGGGAC AGTTTCACGAGTACCAGGAGAGCACAATTGGAGCGGCCTTCCTCACACAGACTGTCTGCCTGGATGA CACAACAGTCAAGTTTGAGATCTGGGACACAGCTGGACAGGAGCGGTATCACAGCCTGGCCCCCATG TACTATCGGGGGGCCCAGGCTGCCATCGTGGTCTATGACATCACCAACACAGATACATTTGCACGGG CCAAGAACTGGGTGAAGGAGCTACAGAGGCAGGCCAGCCCCAACATCGTCATTGCACTCGCGGGTAA		

292

	CAAGGCAGACCTGGCCAGCAAGAGAGCCGTGGAATTCCAGGAAGCACAAGCCTATGCAGACGACAAC AGTTTGCTGTTCATGGAGACATCAGCAAAGACTGCAATGAACGTGAACGAAATCTTCATGGCAATAG CTAAGAAGCTTCCCAAGAACGAGCCCCAGAATGCAACTGGTGCTCCAGGCCGAAACCGAGGTGTGGA CCTCCAGGAGAACAACCCAGCCAGCCGGAGCCAGTGCTGCAGCAACTGAGCCCCCCTTGCCTGCCCG CTGCCCCCGCCTCCTCCGCCTGAATGACCCGACTGGAATCCACTCTAACCAATCGCACTTAACGACT CGGGCCACCACTGGGGGGGCAGGGGGAGGGGTCCACCATGATTTCTCCATATAATTTTGATCATAGG CCGGAGTGAGTCATTCCACCTG		
	ORF Start: ATG at 136		ORF Stop: TGA at 784

	SEQ ID NO: 218	216 aa	MW at 23567.4kD
NOV50a, CG57284-01 Protein Sequence	MAGRGGAARRPNGPAAGNKICQFKLVLLGESAVGKSSLVLRFVKGQFHEYQESTIGAAFLTQTVCLDD TTVKFEIWDTAGQERYHSLAPMYYRGAQAAIVVYDITNTDTFARAKNWVKELQRQASPNIVIALAGN KADLASKRAVEFQEAQAYADDNSLLFMETSAKTAMNVNEIFMAIAKKLPKNEPQNATGAPGRNRGVD LQENNPASRSQCCSN		

	SEQ ID NO: 219	747 bp	
NOV50b, CG57284-03 DNA Sequence	CCACTAAGTGCCTCTTTGCATAGCACCAGTCCCCACCCGCACGCTCTCTGGACCACTACAGCTGGAC GGGCAATGGCGGGTCGGGGAGGCGCAGCACGACCCAATGGACCAGCTGCTGGGGAACAAGATCTGTCA ATTTAAGCTGGTTCTGCTGGGGGAGTCTGCGGTAGGCAAATCCAGCCTCGTCCTCCGCTTTGTCAAG GGACAGTTTCACGAGTACCAGGAGAGCACAATTGGAGCGGCCTTCCTCACACAGACTGTCTGCCTGG ATGACACAACAGTCAAGTTTGAGATCTGGGACACAGCTGGACAGGAGCGGTATCACAGCCTGGCCCC CATGTACTATCGGGGGGCCCAGGCTGCCATCGTGGTCTATGACATCACCAACATCGTCATTGCGCTC GCGGGTAACAAGGCAGACCTGGCCAGCAAGAGAGCCGTGGAATTCCAGGAAGCACAAGCCTATGCAG ACGACAACAGTTTGCTGTTCATGGAGACATCAGCAAAGACTGCAATGAACGTGAACGAAATCTTCAT GGCAATAGCTAAGAAGCTTCCCAAGAACGAGCCCCAGAATGCAACTGGTGCTCCAGGCCGAAACCGA GGTGTGGACCTCCAGGAGAACAACCCAGCCAGCCGGAGCCAGTGCTGCAGCAACTGAGCCCCCCTTG CCTGCCCGCTGCCCCCGCCTCCTCCGCCTGAATGACCCGACTGGAATCCACTCTAACCAATCGCACT TAACGACTCG		
	ORF Start: ATG at 73		ORF Stop: TGA at 658

	SEQ ID NO: 220	195 aa	MW at 21039.6kD
NOV50b, CG57284-03 Protein Sequence	MAGRGGAARPNGPAAGNKICQFKLVLLGESAVGKSSLVLRFVKGQFHEYQESTIGAAFLTQTVCLDD TTVKFEIWDTAGQERYHSLAPMYYRGAQAAIVVYDITNIVIALAGNKADLASKRAVEFQEAQAYADD NSLLFMETSAKTAMNVNEIFMAIAKKLPKNEPQNATGAPGRNRGVDLQENNPASRSQCCSN		

	SEQ ID NO: 221	819 bp	
NOV50c, CG57284-02 DNA Sequence	AATCGCCTTCCACTAAGTGCCTCTTTGCATAGCACCAGTCCCCACCCGCACGCTCTCTGGACCACTA CAGCTGGACGGGCAATGGCGGGTCGGGGAGGCGCAGCACGACCCAATGGACCAGCTGCTGGGGAACAA GATCTGTCAATTTAAGCTGGTTCTGCTGGGGGAGTCTGCGGTAGGCAAATCCAGCCTCGTCCTCCGC TTTGTCAAGGGACAGTTTCACGAGTACCAGGAGAGCACAATTGGAGCGGCCTTCCTCACACAGACTG TCTGCCTGGATGACACAACAGTCAAGTTTGAGATCTGGGACACAGCTGGACAGGAGCGGTATCACAG CCTGGCCCCCATGTACTATCGGGGGGCCCAGGCTGCCATCGTGGTCTATGACATCACCAACACAGAT ACATTTGCACGGGCCAAGAACTGGGTGAAGGAGCTACAGAGGCAGGCCAGCCCCAACATCGTCATTG CACTCGCGGGTAACAAGGCAGACCTGGCCAGCAAGAGAGCCGTGGAATTCCAGGAAGCACAAGCCTA TGCAGACGACAACAGTTTGCTGTTCATGGAGACATCAGCAAAGACTGCAATGAACGTGAACGAAATC		

	TTCATGGCAATAGCTAAGAAGCTTCCCAAGAACGAGCCCCAGAATGCAACTGGTGCTCCAGGCCGAA ACCGAGGTGTGGACCTCCAGGAGAACAACCCAGCCAGCCGGAGCCAGTGCTGCAGCAACTGAGCCCC CCTTGCCTGCCCGCTGCCCCCGCCTCCTCCGCCTGAATGACCCGACTGGAATCCACTCTAACCAATC GCACTTAACGACTCG
ORF Start: ATG at 82		ORF Stop: TGA at 730

	SEQ ID NO: 222	216 aa	MW at 23482.3kD
NOV50c, CG57284-02 Protein Sequence	MAGRGGAARPNGPAAGNKICQFKLVLLGESAVGKSSLVLRFVKGQFHEYQESTIGAAFLTQTVCLDD TTVKFEIWDTAGQERYHSLAPMYYRGAQAAIVVYDITNTDTFARAKNWVKELQRQASPNIVIALAGN KADLASKRAVEFQEAQAYADDNSLLFMETSAKTAMNVNEIFMAIAKKLPKNEPQNATGAPGRNRGVD LQENNPASRSQCCSN

Sequence comparison of the above protein sequences yields the following sequence
relationships shown in Table 50B.

10

Table 50B. Comparison of NOV50a against NOV50b and NOV50c.		
Protein Sequence	NOV50a Residues/ Match Residues	Identities/ Similarities for the Matched Region
NOV50b	18..216 18..195	178/199 (89%) 178/199 (89%)
NOV50c	18..216 18..216	199/199 (100%) 199/199 (100%)

Further analysis of the NOV50a protein yielded the following properties shown in
15   Table 50C.

Table 50C. Protein Sequence Properties NOV50a	
PSort analysis:	0.6500 probability located in cytoplasm; 0.2189 probability located in lysosome (lumen); 0.1000 probability located in mitochondrial matrix space; 0.0000 probability located in endoplasmic reticulum (membrane)
SignalP analysis:	No Known Signal Sequence Predicted

20   A search of the NOV50a protein against the Geneseq database, a proprietary
database that contains sequences published in patents and patent publication, yielded
several homologous proteins shown in Table 50D.

Table 50D. Geneseq Results for NOV50a

Geneseq Identifier	Protein/Organism/Length [Patent #, Date]	NOV50a Residues/ Match Residues	Identities/ Similarities for the Matched Region	Expect Value
AAM79225	Human protein SEQ ID NO 1887 - Homo sapiens, 215 aa. [WO200157190-A2, 09-AUG-2001]	9..216 8..215	179/208 (86%) 194/208 (93%)	e-101
AAY56173	Human Wnt-1 amino acid sequence - Homo sapiens, 215 aa. [CA2200794-A, 24-SEP-1998]	9..216 8..215	179/208 (86%) 194/208 (93%)	e-101
AAB28187	Human RAS-relates protein RAB-5A - Homo sapiens, 193 aa. [WO200052165-A2, 08-SEP-2000]	1..197 1..192	178/197 (90%) 186/197 (94%)	9e-97
AAM80209	Human protein SEQ ID NO 3855 - Homo sapiens, 255 aa. [WO200157190-A2, 09-AUG-2001]	9..216 47..255	172/209 (82%) 189/209 (90%)	1e-95
ABB60036	Drosophila melanogaster polypeptide SEQ ID NO 6900 - Drosophila melanogaster, 219 aa. [WO200171042-A2, 27-SEP-2001]	2..214 11..218	159/213 (74%) 177/213 (82%)	8e-85

In a BLAST search of public sequence datbases, the NOV50a protein was found to have homology to the proteins shown in the BLASTP data in Table 50E.

Table 50E. Public BLASTP Results for NOV50a

Protein Accession Number	Protein/Organism/Length	NOV50a Residues/ Match Residues	Identities/ Similarities for the Matched Portion	Expect Value
P51148	Ras-related protein Rab-5C (RAB5L) (L1880) - Homo sapiens (Human), 216 aa.	1..216 1..216	216/216 (100%) 216/216 (100%)	e-122

AAM21086	Small GTP binding protein RAB5C - Homo sapiens (Human), 216 aa.	1..216 1..216	215/216 (99%) 215/216 (99%)	e-121
Q8R1V8	Hypothetical 23.4 kDa protein - Mus musculus (Mouse), 216 aa.	1..216 1..216	212/216 (98%) 213/216 (98%)	e-119
P51147	Ras-related protein Rab-5C - Canis familiaris (Dog), 216 aa.	1..216 1..216	212/216 (98%) 213/216 (98%)	e-119
Q98932	Rab5C-like protein - Gallus gallus (Chicken), 216 aa.	1..216 1..216	203/216 (93%) 208/216 (95%)	e-114

PFam analysis predicts that the NOV50a protein contains the domains shown in the Table 50F.

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Table 50F. Domain Analysis of NOV50a			
Pfam Domain	**NOV50a Match Region**	**Identities/ Similarities for the Matched Region**	**Expect Value**
arf	4..185	40/198 (20%) 105/198 (53%)	0.0018
ras	23..216	90/209 (43%) 181/209 (87%)	3.1e-104

Example 51.

10   The NOV51 clone was analyzed, and the nucleotide and encoded polypeptide sequences are shown in Table 51A.

Table 51A. NOV51 Sequence Analysis			
	SEQ ID NO: 223	4826 bp	
NOV51a, CG57308-01 DNA Sequence	AGCTGAGCCCGAGCCCAGACCGCGCCCGCGCCGCCATGCCCCTGGCCTTCTGCGGCAGCGAGAACCA CTCGGCCGCCTACCGGGTGGACCAGGGGGTCCTCAACAACGGCTGCTTTGTGGACGCGCTCAACGTG GTGCCGCACGTCTTCCTACTCTTCATCACCTTCCCCATCCTCTTCATTGGATGGGGAAGTCAGAGCT CCAAGGTGCACATCCACCACAGCACATGGCTTCATTTCCCTGGGCACAACCTGCGGTGGATCCTGAC CTTCATGCTGCTCTTCGTCCTGGTGTGTGAGATTGCAGAGGGCATCCTGTCTGATGGGGTGACCGAA TCCCACCATCTGCACCTGTACATGCCAGCCGGGATGGCGTTCATGGCTGCTGTCACCTCCGTGGTCT ACTATCACAACATCGAGACTTCCAACTTCCCCAAGCTGCTAATTGCCCTGCTGGTGTATTGGACCCT GGCCTTCATCACCAAGACCATCAAGTTTGTCAAGTTCTTGGACCACGCCATCGGCTTCTCGCAGCTA CGCTTCTGCCTCACAGGGCTGCTGGTGATCCTCTATGGGATGCTGCTCCTCGTGGAGGTCAATGTCA TCAGGGTGAGGAGATACATCTTCTTCAAGACACCGAGGGAGGTGAAGCCTCCCGAGGACCTGCAAGA CCTGGGGGTACGCTTCCTGCAGCCCTTCGTGAATCTGCTGTCCAAAGGCACCTACTGGTGGATGAAC GCCTTCATCAAGACTGCCCACAAGAAGCCCATCGACTTGCGAGCCATCGGGAAGCTGCCCATCGCCA TGAGGGCCCTCACCAACTACCAACGGCTCTGCGAGGCCTTTGACGCCCAGGTGCGGAAGGACATTCA

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GGGCACTCAAGGTGCCCGGGCCATCTGGCAGGCACTCAGCCATGCCTTCGGGGAGGCGCCTGGTCCTC
AGCAGCACTTTCCGCATCTTGGCCGACCTGCTGGGCTTCGCCGGGCCACTGTGCATCTTTGGGATCG
TGGACCACCTTGGGAAGGAGAACGACGTCTTCCAGCCCAAGACACAATTTCTCGGGGTTTACTTTGT
CTCATCCCAAGAGTTCCTTGCCAATGCCTACGTCTTAGCTGTGCTTCTGTTCCTTGCCCTCCTACTG
CAAAGGACATTTCTGCAAGCATCCTACTATGTGGCCATTGAAACTGGAATTAACTTGAGAGGAGCAA
TACAGACCAAGATTTACAATAAAATTATGCACCTGTCCACCTCCAACCTGTCCATGGGAGAAATGAC
TGCTGGACAGATCTGTAATCTGGTTGCCATCGACACCAATCAGCTCATGTGGTTTTTCTTCTTGTGC
CCAAACCTCTGGGCTATGCCAGTACAGATCATTGTGGGTGTGATTCTCCTCTACTACATACTCGGAG
TCAGTGCCTTAATTGGAGCAGCTGTCATCATTCTACTGGCTCCTGTCCAGTACTTCGTGGCCACCAA
GCTGTCTCAGGCCCAGCGGAGCACACTGGAGTATTCCAATGAGCGGCTGAAGCAGACCAACGAGATG
CTCCGCGGCATCAAGCTGCTGAAGCTGTACGCCTGGGAGAACATCTTCCGCACGCGGGTGGAGACGA
CCCGCAGGAAGGAGATGACCAGCCTCAGGGCCTTTGCCATCTATACCTCCATCTCCATTTTCATGAA
CACGGCCATCCCCATTGCAGCTGTCCTCATAACTTTCGTGGGCCATGTCAGCTTCTTCAAAGAGGCC
GACTTCTCGCCCTCCGTGGCCTTTGCCTCCCTCTCCCTCTTCCATATCTTGGTCACACCGCTGTTCC
TGCTGTCCAGTGTGGTCCGATCTACCGTCAAAGCTCTAGTGAGCGTGCAAAAGCTAAGCGAGTTCCT
GTCCAGTGCAGAGATCCGTGAGGAGCAGTGTGCCCCCCATGAGCCCACACCTCAGGGCCCAGCCAGC
AAGTACCAGGCGGTGCCCCTCAGGGTTGTGAACCGCAAGCGTCCAGCCCGGGAGGATTGTCGGGGCC
TCACCGGCCCACTGCAGAGCCTGGTCCCCAGTGCAGATGGCGATGCTGACAACTGCTGTGTCCAGAT
CATGGGGAGGCTACTTCACGTGGACCCCAGATGGAATCCCCACACTGTCCAACATCACCATTCGTATC
CCCCGAGGGCCAGCTGACTATGATCGTGGGGCAGGTGGGCTGCGGCAAGTCCTCGCTCCTTCTAGCCG
CACTGGGGGAGATGCAGAAGGTCTCAGGGGCTGTCTTCTGGAGCAGCCTTCCTGACAGCGAGATAGG
AGAGGACCCCAGCCCAGAGCGGGAGACAGCGACCGACTTGGATATCAGGAAGAGAGGCCCCGTGGCC
TATGCTTCGCAGAAACCATGGCTGCTAAATGCCACTGTGGAGGAGAACATCATCTTTGAGAGTCCCT
TCAACAAACAACGGTACAAGATGGTCATTGAAGCCTGCTCTCTGCAGCCAGACATCGACATCCTGCC
CCATGGAGACCAGACCCTGACCAGATTGGGGAACGGGGCATCAACCTGTCTGGTGGTCAACGCCAGCGAATC
AGTGTGGCCCGAGCCCTCTACCAGCACGCCAACGTTGTCTTCTTGGATGACCCCCTTCTCAGCTCTGG
ATATCCATCTGAGTGACCACTTAATGCAGGCCGGCATCCTTGAGCTGCTCCGGGACGACAAGAGGAC
AGTGGTCTTAGTGACCCACAAGCTACAGTACCTGCCCCATGCAGACTGGATCATTGCCATGAAGGAT
GGCACCATCCAGAGGGAGGGTACCCTCAAGGACTTCCAGAGGGTCTGAATGCCAGCTCTTTGAGCACT
GGAAGACCCTCATGAACCGACAGGACCAAGAGCTGGAGAAGGAGACTGTCACAGAGAGAAAAGCCAC
AGAGCCCACCCCAGGGCCTATCTCGTGCCATGTCCTCGAGGGATGGCCTTCTGCAGGATGAGGAAGAG
GAGGAAGAGGAGGCCAGCTGAGAGCGAGGAGGATGACAACCTGTCGTCCATGCTGCACCAGCGTGCTG
AGATCCCATGGCGAGCCTGCGCCAAGTACCTGTCCTCCGCCGGCATCCTGCTCCTGTCGTTGCTGGT
CTTCTCACAGCTGCTCAAGCACATGGTCCTGGTGGCCATCGACTACTGGCTGGCCAAGTGGACCGAC
AGCGCCCTGACCCCTGACCCCCTGCAGCCAGGAACTGCTCCCTCAGCCAGGAGTGCACCCTCGACCAGA
CTGTCTATGCCATGGTGTTCACGGTGCTCTGCAGCCTGGGCATTGTGCTGCTGTGCCTCGTCACGTCTGT
CACTGTGGAGTGGACAGGGCTGAAGGTGGCCAAGAGACTGCACCGCAGCCTGCTAAACCGGATCATC
CTAGCCCCCATGAGGTTTTTTTGAGACCACGCCCCTTGGGAGCATCCTGAACAGATTTTCATCTGACT
GTAACACCATCGACCAGCACATCCCATCCACGCTGGAGTGCCTGAGCCGCTCCACCCTGCTCTGTGT
CTCAGCCCTGGCCGTCATCTCCTATGTCACACCTGTGTTCCTCGTGGCCCTCTTGCCCCTGGCCATC
GTGTGCTACTTCATCCAGAAGTACTTCCGGGTGGCGTCCAGGGACCTGCAGCAGCTGGATGACACCA
CCCAGCTTCCACTTCTCTCTCACACTTTGCCGAAACCGTAGAAGGACTCACCACCATCCGGGCCTTCAG
GTATGAGGCCCGGTTCCAGCAGAAGCTTCTCGAATACACAGACTCCAACAACATTGCTTCCCTCTTC
CTCACAGCTGCCAACAGATGGCTGGAAGTCCGAATGGAGTACATCGGTGCATGTGTGGTGCTCATCG
CAGCGGTGACCTCCATCTCCAACTCCCTGCACAGGGAGCTCTCTGCTGGCCTGGTGGGCCTGGGCCT
TACCTACGCCCTAATGGTCTCCAACTACCTCAACTGGATGGTGAGGAACCTGGCAGACATGGAGCTC
CAGCTGGGGGCTGTGAAGCGCATCCATGGGCTCCTGAAAACCGAGGCAGAGAGCTACGAGGGGCTCC
TGGCACCATCGCTGATCCCAAAGAACTGGCCAGACCAAGGGAAGATCCAGATCCAGAACCTGAGCGT
GCGCTACGACAGCTCCCTGAAGCCCGATGTGCTGAAGCACGTCAATGCCCTCATCTCCCCTGGACAGAAG
ATCGGGATCTGCGGCCGCACCGGCAGTGGGAAGTCCTCCTTCTCTCTTGCCTTCTTCCGCATGGTGG
ACACGTTCGAAGGGCACATCATCATTGATGGCATTGACATCGCCAAACTGCCGCTGCACACCCTGCG
CTCACGCCTCTCCATCATCCTGCAGGACCCCGTCCTCTTCAGCGGCACCATCCGATTTAACCTGGAC
CCTGAGGAGGAAGTGCTCAGATAGCACACTGTGGGAGGCCCTGGAAATCGCCCAGCTGAAGCTGGTGG
TGAAGGCACTGCCAGGAGGCCTCGATGCCATCATCACAGAAGGCGGGGGAGAATTTCAGCCAGGGACA
GAGGCAGCTGTTCTGCCTGGCCCGGGCCTTCGTGAGGAAGACCAGCATCTTCATCATGGACGAGGCC
ACGGCTTCCATTGACATGGCCACGGAAAACATCCTCCAAAAGGTGGTGATGACAGCCTTCGCAGACC
GCACTGTGGTCACCATCGCGCATCGAGTGCACACCATCCTGAGTGCAGACCTGGTGATCGTCCTGAA
GCGGGGTGCCATCCTTGAGTTCGATAAGCCAGAGAAGCTGCTCAGCCGGAAGGACAGCGTCTTCGCC
TCCTTCGTCCGTGCAGACAAGTGACCTGCCAGAGCCCAAGTGCCATCCCACATTCGGACCCTGCCCA
TA
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ORF Start: ATG at 36		ORF Stop: TGA at 4779

	SEQ ID NO: 224	1581 aa	MW at 177005.9kD
NOV51a, CG57308-01 Protein Sequence	MPLAFCGSENHSAAYRVDQGVLNNGCFVDALNVVPHVFLLFITPPILFIGWGSQSSKVHIHHSTWLH FPGHNLRWILTFMLLFVLVCEIAEGILSDGVTESHHLHLYMPAGMAFMAAVTSVVYYHNIETSNFPK LLIALLVYWTLAFITKTIKFVKFLDHAIGFSQLRFCLTGLLVILYGMLLLVEVNVIRVRRYIFFKTP REVKPPEDLQDLGVRFLQPFVNLLSKGTYWWMNAFIKTAHKKPIDLRAIGKLPIAMRALTNYQRLCE AFDAQVRKDIQGTQGARAIWQALSHAFGRRLVLSSTFRILADLLGFAGPLCIFGIVDHLGKENDVFQ		

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| | PKTQFLGVYFVSSQEFLANAYVLAVLLFLALLLQRTFLQASYYVAIETGINLRGAIQTKIYNKIMHL
STSNLSMGEMTAGQICNLVAIDTNQLMWFFFLCPNLWAMPVQIIVGVILLYYILGVSALIGAAVIIL
LAPVQYFVATKLSQAQRSTLEYSNERLKQTNEMLRGIKLLKLYAWENIFRTRVETTRRKEMTSLRAF
AIYTSISIFMNTAIPIAAVLITFVGHVSFFKEADFSPSVAFASLSLFHILVTPLFLLSSVVRSTVKA
LVSVQKLSEFLSSAEIREEQCAPHEPTPQGPASKYQAVPLRVVNRKRPAREDCRGLTGPLQSLVPSA
DGDADNCCVQIMGGYFTWTPDGIPTLSNITIRIPRGQLTMIVGQVGCGKSSLLLAALGEMQKVSGAV
FWSSLPDSEIGEDPSPERETATDLDIRKRGPVAYASQKPWLLNATVEENIIPESPPFNKQRYKMVIEA
CSLQPDIDILPHGDQTQIGERGINLSGGQRQRISVARALYQHANVVFLDDPPSALDIHLSDHLMQAG
ILELLRDDKRTVVLVTHKLQYLPHADWIIAMKDGTIQREGTLKDFQRSECQLFEHWKTLMNRQDQEL
EKETVTERKATEPPQGLSRAMSSRDGLLQDEEEEEEEAAESEEDDNLSSMLHQRAEIPWRACAKYLS
SAGILLLSLLVFSQLLKHMVLVAIDYWLAKWTDSALTLTPAARNCSLSQECTLDQTVYAMVFTVLCS
LGIVLCLVTSVTVEWTGLKVAKRLHRSLLNRIILAPMRFFETTPLGSILNRFSSDCNTIDQHIPSTL
ECLSRSTLLCVSALAVISYVTPVFLVALLPLAIVCYFIQKYFRVASRDLQQLDDTTQLPLLSHFAET
VEGLTTIRAFRYEARFQQKLLEYTDSNNIASLFLTAANRWLEVRMEYIGACVVLIAAVTSISNSLHR
ELSAGLVGLGLTYALMVSNYLNWMVRNLADMELQLGAVKRIHGLLKTEAESYEGLLAPSLIPKNWPD
QGKIQIQNLSVRYDSSLKPVLKHVNALISPGQKIGICGRTGSGKSSFSLAFFRMVDTFEGHIIDGI
DIAKLPLHTLRSRLSIILQDPVLFSGTIRFNLDPERKCSDSTLWEALEIAQLKLVVKALPGGLDAII
TEGGENFSQGQRQLFCLARAFVRKTSIFIMDEATASIDMATENILQKVVMTAFADRTVVTIAHRVHT
ILSADLVIVLKRGAILEFDKPEKLLSRKDSVFASFVRADK |

	SEQ ID NO: 225	4745 bp	
NOV51b, CG57308-02 DNA Sequence	CGGGGCCCGGGGGGCGGGGGCCTGACGGCCGGGCCGGGCCGGGCGGCGGAGCTGCAAGGGACAGAGGCGCGG CAGGCGCGCGGAGCCAGCGGAGCCAGCTGAGCCCGAGCCCAGCCCGCGCCCCGCGCCGCCATGCCCCT GGCCTTCTGCGGCAGCGAGAACCACTCGGCCGCCTACCGGGTGGACCAGGGGGTCCTCAACAACGGC TGCTTTGTGGACGCGCTCAACGTGGTGCCGCACGTCTTCCTACTCTTCATCACCTTCCCCATCCTCT TCATTGGATGGGGAAGTCAGAGCTCCAAGGTGCACATCCACCACAGCACATGGCTTCATTTCCCCGG GCACAACCTGCGGTGGATCCTGACCTTCATGCTGCTCTTCGTCCTGGTGTGTGAGATTGCAGAGGGC ATCCTGTCTGATGGGGTGACCGAATCCCACCATCTGCACCTGTACATGCCAGCCGGGATGGCCGTTCA TGGCTGCTGTCACCTCCGTGGTCTATATCACAACATCGAGACTTCCAACTTCCCCAAGCTGCTAAT TGCCCTGCTGGTGTATTGGACCCTGGCCTTCATCACCAAGACCATCAAGTTTGTCAAGCTCTTGGAC CACGCCATCGGCTTCTCGCAGCTACGCTTCTGCCTCACAGGGCTGCTGGTGATCCTCTATGGGATGC TGCTCCTCGTGGGAGGTCAATGTCATCAGGGTGAGGAGATACATCTTCTTCAAGACACCGAGGGAGGT GAAGCCTCCCGAGGACCTGCAAGACCTGGGGGTACGCTTCCTGCAGCCCTTCGTGAATCTGCCGTCC AAAGGCACCTACTGGTGGATGAACGCCTTCATCAAGACTGCCCACAAGAAGCCCATCGACTTGCGAG CCATCGGGAAGCTGCCCATCGTTATGAGGGCCCTCACCAACTACCAACGGCTCTGCGAGGCCTTTGA CGCCCAGGTGCCGGAAGGACATTCAGGGCACTCAAGGTGCCCGGGCCCATCTGGCAGGCACTCAGCCAT GCCTTCGGGAGGCGCCTGGTCCTCAGCAGCACTTTCCGCATCTTGGCCGACCTGCTGGGCTTCGCCG GGCCACTGTGCATCTTTGGGATCGTGGACCACCTTGGGAAGGAGAACGACGTCTTCCAGCCCAAGAC ACAATTTCTCGGGGTTTACTTTGTCTCATCCCAAGAGTTCCTTGCCAATGCCTACGTCTTAGCTGTG CTTCTGTTCCTTGCCCTCCTACTGCAAAGGACATTTCTGCAAGCATCCTACTATGTGGCCATTGAAA CTGGAATTAACTTGAGAGGAGCAATACAGACCAAGATTTACAATAAAATTATGCACCTGTCCACCTC CAACCTGTCCATGGGAGAAATGACTGCTGGACAGATCTGTAATCTGGTTGCCATCGACACCAATCAG CTCATGTGGTTTTTCTTCTTGTGCCAAACCTCTGGGCTATGCCAGTACAGATCATTGTGGGTGTGA TTCTCCTCTACTACATACTCGGAGTCAGTGCCTTAATTGGAGCAGCTGTCATCATTCTACTGGCTCC TGTCCAGTACTTCGTGGCCACCAAGCTGTCTCAGGCCCAGCGGAGCACACTGGAGTATTCCAATGAG CGGCTGAAGCAGACCAACGAGATGCTCCGCGGCATCAAGCTGCTGAAGCTGTACGCCTGGGAGAACA TCTTCCGCACGCGGGTGGAGACGACCCGCAGGAAGGAGATGACCAGCCTCAGGGCCTTTGCCATCTA TACCTCCATCTCCATTTTCATGAACACGGCCATCCCCATTGCAGCTGTCCTCATAACTTTCGTGGGC CATGTCAGCTTCTTCAAAGAGGCCGACTTCTCGCCCTCCGTGGCCTTTGCCTCCCTCTCCCTCTTCC ATATCTTGGTCACACCGCTGTTCCTGCTGTCCAGTGTGGTCCGATCTACCGTCAAAGCTCTAGTGAG CGTGCAAAAGCTAAGCGAGTTCCTGTCCAGTGCAGAGATCCGTGAGGAGCAGTGTGCCCCCCATGAG CCCACACCTCAGGGCCCAGCCAGCAAGTACCAGGCGGTGCCCCTCAGGGTTGTGAACCGCAAGCGTC CAGCCGGGAGGATTGTCGGGGCCTCACCGGCCCACTGCAGAGCCTGGTCCCCAGTGCAGATGGCGA TGCTGACAACTGCTGTGTCCAGATCATGGGAGGCTACTTCACGTGGACCCCAGATGGAATCCCCACA CTGTCCAACATCACCATTCGTATCCCCGAGGCCAGCTGACTATGATCGTGGGGCAGGTGGGCTGCG GCAAGTCCTCGCTCCTTCTAGCCGCACTGGGGGGAGATGCAGAAGGTCTCAGGGGCTGTCTTCTGGAG CAGCCTTCCTGACAGCGAGATAGGAGGAGGACCCCAGCCCAGAGCGGGAGACAGCGACCGACTTGGAT ATCAGGAAGAGAGGCCCCGTGGCCTATGCTTCGCAGAAACCATGGCTGCTAAATGCCACTGTGGAGG AGAACATCATCTTTGAGAGTCCCTTCAACAAACAACGGTACAAGATGGTCATTGAAGCCTGCTCTCT GCAGCCAGACATCGACATCCTGCCCCATGGAGACCAGACCCAGATTGGGGAACGGGGCATCAACCTG TCTGGTGGTCAACGCCAGCGAATCAGTGTGGCCCGAGCCCTCTACCAGCACGCCAACGTTGTCTTCT TGGATGACCCCTTCTCAGCTCTGGATATCCATCTGAGTGACCACTTAATGCAGGCCGGCATCCTTGA GCTGCTCCGGGACGACAAGAGGACAGTGGTCTTAGTGACCCACAAGCTACAGTACCTGCCCCATGCA GACTGGATCATTGCCATGAAGGATGGCACCATCCAGAGGGAGGGTACCCTCAAGGACTTCCAGAGGT CTGAATGCCAGCTCTTTGAGCACTGGAAGACCCTCATGAACCGACAGGACCAAGAGCTGGAGAAGGA GACTGTCACAGAGAGAAAAGCCACAGAGCCACCCCAGGGCCTATCTCGTGCCATGTCCTCGAGGGAT GGCCTTCTGCAGGATGAGGAAGAGGAGGAAGAGGAGGCAGCTGAGAGCGAGGAGGATGACAACCTGT CGTCCATGCTGCACCAGCGTGCTGAGATCCCATGGCGAGCCTGCGCCAAGTACCTGTCCTCCGCCGG CATCCTGCTCCTGTCGTTGCTGGTCTTCTCACAGCTGCTCAAGCACATGGTCCTGGTGGCCATCGAC

	TACTGGCTGGCCAAGTGGACCGACAGCGCCCTGACCCTGACCCCTGCACCCTGCAGCCAGGAACTGCTCCCTCA		
	GCCAGGAGTGCACCCTCGACCAGACTGTCTATGCCATGGTGTTCACGGTGCTCTGCAGCCTGGGCAT		
	TGTGCTGTGCCTCGTCACGTCTGTCACTGTGGAGTGGACAGGGCTGAAGGTGGCCAAGAGACTGCAC		
	CGCAGCCTGCTAAACCGGATCATCCTAGCCCCCATGAGGTTTTTTGAGACCACGCCCCTTGGGAGCA		
	TCCTGAACAGATTTTCATCTGACTGTAACACCATCGACCAGCACATCCCATCCACGCTGGAGTGCCT		
	GAGCCGCTCCACCCTGCTCTGTGTCTCAGCCCTGGCCGTCATCTCCTATGTCACACCTGTGTTCCTC		
	GTGGCCCTCTTGCCCCTGGCCATCGTGTGCTACTTCATCCAGAAGTACTTCCGGGTGGCGTCCAGGG		
	ACCTGCAGCAGCTGGATGACACCACCCAGCTTCCACTTCTCTCACACTTTGCCGAAACCGTAGAAGG		
	ACTCACCACCATCCGGGCCTTCAGGTATGAGGCCCGGTTCCAGCAGAAGCTTCTCGAATACACAGAC		
	TCCAACAACATTGCTTCCCTCTTCCTCACAGCTGCCAACAGATGGCTGGAAGTCCGAATGGAGTACA		
	TCGGTGCATGTGTGGTGCTCATCGCCAGCGGTGACCTCCATCTCCAACTCCCTGCACAGGGAGCTCTC		
	TGCTGGCCTGGTGGGCCTGGGCCTTACCTACGCCCTAATGGTCTCCAACTACCTCAACTGGATGGTG		
	AGGAACCTGGCAGACATGGAGCTCCAGCTGGGGGCTGTGAAGCGCATCCATGGGCTCCTGAAAACCG		
	AGGCAGAGAGCTACGAGGGGCTCCTGGCACCATCGCTGATCCCAAAGAACTGGCCAGACCAAGGGAA		
	GATCCAGATCCAGAACCTGAGCGTGCGCGATACGACAGCTCCCTGAAGCCGGTGCTGAAGCACGTCAAT		
	GCCCTCATCTCCCCTGGACAGAAGATCGGGATCTGCGGCCGCACCGGCAGTGGGAAGTCCTCCTTCT		
	CTCTTGCCTTCTTCCGCATGGTGGACACGTTCGAAGGGCACATCATCACAGAAGGCGGGGAGAATTT		
	CAGCCAGGGACAGAGGCAGCTGTTCTGCCTGGCCCGGGCCTTCGTGAGGAAGACCAGCATCTTCATC		
	ATGGACGAGGCCACGGCTTCCATTGACATGGCCACGGAAAACATCCTCCAAAAGGTGGTGATGACAG		
	CCTTCGCAGACCGCACTGTGGTCACCATCGCGCATCGAGTGCACACCATCCTGAGTGCAGACCTGGT		
	GATCGTCCTGAAGCGGGGTGCCATCCTTGAGTTCGATAAGCCAGAGAAGCTGCTCAGCCGGAAGGAC		
	AGCGTCTTCGCCTCCTTCGTCCGTGCAGACAAGTGACCTGCCAGAGCCCAAGTGCCATCCCACATTC		
	GGACCCTGCCCATACCCCTGCCTGGGTTTTCTAACTGTAAATCACTTGTAAATAA		
ORF Start: ATG at 127			ORF Stop: TGA at 4657

	SEQ ID NO: 226	1510 aa	MW at 169179.9kD
NOV51b, CG57308-02 Protein Sequence	MPLAFCGSENHSAAYRVDQGVLNNGCFVDALNVVPHVFLLFITFPILFIGWGSQSSKVHIHHSTWLH FPGHNLRWILTFMLLFVLVCEIAEGILSDGVTESHHLHLYMPAGMAFMAAVTSVVYYHNIETSNFPK LLIALLVYWTLAFITKTIKFVKLLDHAIGFSQLRFCLTGLLVILYGMLLLVEVNVIRVRRYIFFKTP REVKPPEDLQDLGVRFLQPFVNLPSKGTYWWMNAFIKTAHKKPIDLRAIGKLPIVMRALTNYQRLCE AFDAQVRKDIQGTQGARAIWQALSHAFGRRLVLSSTFRILADLLGFAGPLCIFGIVDHLGKENDVFQ PKTQFLGVYFVSSQEFLANAYVLAVLLFLALLLQRTFLQASYYVAIETGINLRGAIQTKIYNKIMHL STSNLSMGEMTAGQICNLVAIDTNQLMWFFFLCPNLWAMPVQIIVGVILLYYILGVSALIGAAVIIL LAPVQYFVATKLSQAQRSTLEYSNERLKQTNEMLRGIKLLKLYAWENIFRTRVETTRRKEMTSLRAF AIYTSISIFMNTAIPIAAVLITFVGHVSFFKEADFSPSVAFASLSLFHILVTPLFLLSSVVRSTVKA LVSVQKLSEFLSSAEIREEQCAPHEPTPQGPASKYQAVPLRVVNRKRPAREDCRGLTGPLQSLVPSA DGDADNCCVQIMGGYFTWTPDGIPTLSNITIRIPRGQLTMIVGQVGCGKSSLLLAALGEMQKVSGAV FWSSLPDSEIGEDPSPERETATDLDIRKRGPVAYASQKPWLLNATVEENIIFESPFNKQRYKMVIEA CSLQPDIDILPHGDQTQIGERGINLSGGQRQRISVARALYQHANVVFLDDPFSALDIHLSDHLMQAG ILELLRDDKRTVVLVTHKLQYLPHADWIIAMKDGTIQREGTLKDFQRSECQLFEHWKTLMNRQDQEL EKETVTERKATEPPQGLSRAMSSRDGLLQDEEEEEEEAAESEEDDNLSSMLHQRAEIPWRACAKYLS SAGILLLSLLVFSQLLKHMVLVAIDYWLAKWTDSALTLTPAARNCSLSQECTLDQTVYAMVFTVLCS LGIVLCLVTSVTVEWTGLKVAKRLHRSLLNRIILAPMRFFETTPLGSILNRFSSDCNTIDQHIPSTL ECLSRSTLLCVSALAVISYVTPVFLVALLPLAIVCYFIQKYFRVASRDLQQLDDTTQLPLLSHFAET VEGLTTIRAFRYEARFQQKLLEYTDSNNIASLFLTAANRWLEVRMEYIGACVVLIAAVTSISNSLHR ELSAGLVGLGLTYALMVSNYLNWMVRNLADMELQLGAVKRIHGLLKTEAESYEGLLAPSLIPKNWPD QGKIQIQNLSVRYDSSLKPVLKHVNALISPGQKIGICGRTGSGKSSFSLAFFRMVDTFEGHIITEGG ENFSQGQRQLFCLARAFVRKTSIFIMDEATASIDMATENILQKVVMTAFADRTVVTIAHRVHTILSA DLVIVLKRGAILEFDKPEKLLSRKDSVFASFVRADK		

Sequence comparison of the above protein sequences yields the following sequence relationships shown in Table 51B.

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Table 51B. Comparison of NOV51a against NOV51b.		
Protein Sequence	NOV51a Residues/ Match Residues	Identities/ Similarities for the Matched Region

| NOV51b | 1..1406 | 1285/1406 (91%) |
| | 1..1406 | 1286/1406 (91%) |

Further analysis of the NOV51a protein yielded the following properties shown in Table 51C.

Table 51C. Protein Sequence Properties NOV51a

| PSort analysis: | 0.8000 probability located in plasma membrane; 0.4000 probability located in Golgi body; 0.3000 probability located in endoplasmic reticulum (membrane); 0.3000 probability located in microbody (peroxisome) |
| SignalP analysis: | Cleavage site between residues 56 and 57 |

A search of the NOV51a protein against the Geneseq database, a proprietary database that contains sequences published in patents and patent publication, yielded several homologous proteins shown in Table 51D.

Table 51D. Geneseq Results for NOV51a

Geneseq Identifier	Protein/Organism/Length [Patent #, Date]	NOV51a Residues/ Match Residues	Identities/ Similarities for the Matched Region	Expect Value
AAW57412	Homo sapiens sulphonylurea receptor - Homo sapiens, 1580 aa. [WO9814571-A1, 09-APR-1998]	1..1581 1..1580	1530/1582 (96%) 1540/1582 (96%)	0.0
AAR77087	Rat sulphonylurea receptor - Rattus sp, 1582 aa. [WO9528411-A1, 26-OCT-1995]	1..1581 1..1582	1477/1582 (93%) 1509/1582 (95%)	0.0
AAR77088	Hamster sulphonylurea receptor - Cricetus sp, 1582 aa. [WO9528411-A1, 26-OCT-1995]	1..1581 1..1582	1469/1582 (92%) 1506/1582 (94%)	0.0
AAR77084	Rat sulphonylurea receptor - Rattus sp, 1498 aa. [WO9528411-A1, 26-OCT-1995]	1..1290 1..1291	1195/1291 (92%) 1223/1291 (94%)	0.0

300

| AAR77085 | Hamster sulphonylurea receptor - Cricetus sp, 1498 aa. [WO9528411-A1, 26-OCT-1995] | 1..1290 1..1291 | 1186/1291 (91%) 1220/1291 (93%) | 0.0 |

In a BLAST search of public sequence datbases, the NOV51a protein was found to have homology to the proteins shown in the BLASTP data in Table 51E.

Table 51E. Public BLASTP Results for NOV51a				
Protein Accession Number	Protein/Organism/Length	NOV51a Residues/ Match Residues	Identities/ Similarities for the Matched Portion	Expect Value
Q09428	Sulfonylurea receptor 1 - Homo sapiens (Human), 1580 aa.	2..1581 1..1580	1579/1580 (99%) 1579/1580 (99%)	0.0
Q09429	Sulfonylurea receptor 1 - Rattus norvegicus (Rat), 1581 aa.	2..1581 1..1581	1512/1582 (95%) 1536/1582 (96%)	0.0
Q09427	Sulfonylurea receptor 1 - Cricetus cricetus (Black-bellied hamster), 1581 aa.	2..1581 1..1581	1498/1582 (94%) 1530/1582 (96%)	0.0
A56248	sulfonylurea receptor - golden hamster, 1582 aa.	1..1581 1..1582	1469/1582 (92%) 1506/1582 (94%)	0.0
Q95J92	Sulphonylurea receptor 2B - Oryctolagus cuniculus (Rabbit), 1549 aa.	1..1580 1..1548	1076/1581 (68%) 1277/1581 (80%)	0.0

PFam analysis predicts that the NOV51a protein contains the domains shown in the Table 51F.

Table 51F. Domain Analysis of NOV51a			
Pfam Domain	NOV51a Match Region	Identities/ Similarities for the Matched Region	Expect Value
ABC_membrane	318..590	53/287 (18%) 212/287 (74%)	3.6e-46

ABC_tran	706..905	55/214 (26%) 154/214 (72%)	1.3e-34
ABC_membrane	1011..1298	58/292 (20%) 222/292 (76%)	2.7e-51
PRK	1374..1391	6/19 (32%) 15/19 (79%)	0.21
ABC_tran	1371..1554	54/199 (27%) 129/199 (65%)	5.7e-36

Example 52.

The NOV52 clone was analyzed, and the nucleotide and encoded polypeptide

5   sequences are shown in Table 52A.

Table 52A. NOV52 Sequence Analysis			
	SEQ ID NO: 227	1404 bp	
NOV52a, CG93659-01 DNA Sequence	ATGGAGTACATGAGCACTGGAAGTGACAATAAAGAAGAGATTGATTTATTAATTAAACATTTAAATG TGTCTGATGTAATAGACATTATGGAAAATCTTTATGCAAGTGAAGAGCCAGCAGTTTATGAACCCAG TCTAATGACCATGTGTCAAGACAGTAATCAAAACGATGAGCGTTCTAAGTCTCTGCTGCTTAGTGGC CAAGAGGTACCATGGTTGTCATCAGTCAGATATGGAACTGTGGAGGATTTGCTTGCTTTTGCAAACC ATATATCCAACACTGCAAAGCATTTTTATGGACAACGACCACAGGAATCTGGAATTTTATTAAACAT GGTCATCACTCCCCAAAATGGACGTTACCAAATAGATTCCGATGTTCTCCTGATCCCCTGGAAGCTG ACTTACAGGAATATTGGTTCTGATTTTATTCCTCGGGGCGCCTTTGGAAAGGTATACTTGGCTCAAG ATATAAAGACGAAGAAAAGAATGGCGTGTAAACTGATCCCAGTAGATCAATTTAAGCCATCTGATGT GGAAATTCAGGCTTGCTTCCGGCACGAGAACATCGCAGAGCTGTATGGCGCAGTCCTGTGGGGTGAA ACTGTCCATCTCTTTATGGAAGCAGGCGAGGGAGGGTCTGTTCTGGAGAAACTGGAGAGCTGTGGAC CAATGAGAGAATTTGAAATTATTTGGGTGACAAAGCATGTTCTCAAGGGACTTGATTTTCTACACTC AAAGAAAGTGATCCATCATGATATTAAACCTAGCAACATTGTTTTCATGTCCACAAAAGCTGTTTTG GTGGATTTTGGCCTAAGTGTTCAAATGACCGAAGATGTCTATTTTCCTAAGGACCTCCGAGGAACAG AGATTTACATGAGCCCAGAGGTCATCCTGTGCAGGGGCCATTCAACCAAAGCAGACATCTACAGCCT GGGGGCCACGCTCATCCACATGCAGACGGGCACCCCACCCTGGGTGAAGCGCTACCCTCGCTCAGCC TATCCCTCCTACCTGTACATAATCCACAAGCAAGCACCTCCACTGGAAGACATTGCAGATGACTGCA GTCCAGGGATGAGAGAGCTGATAGAAGCTTCCCTGGAGAGAAACCCCAATCACCGCCCAAGAGCCGC AGACCTACTAAAACATGAGGCCCTGAACCCGCCCAGAGAGGATCAGCCACGCTGTACGAGTCTGGAC TCTGCCCTCTTGGAGCGCAAGAGGCTGCTGAGTAGGAAGGAGCTGGAACTTCCTGAGAACATTGCTG ATTCTTCGTGCACAGGAAGCACCGAGGAATCTGAGATGCTCAAGAGGCAACGCTCTCTCTACATCGA CCTCGGCGCTCTGGCTGGCTACTTCAATCTTGTTCGGGGACCACCAACGCTTGAATATGGCTGA		
	ORF Start: ATG at 1		ORF Stop: TGA at 1402

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	SEQ ID NO: 228	467 aa	MW at 52896.9kD
NOV52a, CG93659-01 Protein Sequence	MEYMSTGSDNKEEIDLLIKHLNVSDVIDIMENLYASEEPAVYEPSLMTMCQDSNQNDERSKSLLLSG QEVPWLSSVRYGTVEDLLAFANHISNTAKHFYGQRPQESGILLNMVITPQNGRYQIDSDVLLIPWKL TYRNIGSDFIPRGAFGKVYLAQDIKTKKRMACKLIPVDQFKPSDVEIQACFRHENIAELYGAVLWGE TVHLFMEAGEGGSVLEKLESCGPMREFEIIWVTKHVLKGLDFLHSKKVIHHDIKPSNIVFMSTKAVL VDFGLSVQMTEDVYFPKDLRGTEIYMSPEVILCRGHSTKADIYSLGATLIHMQTGTPPWVKRYPRSA YPSYLYIIHKQAPPLEDIADDCSPGMRELIEASLERNPNHRPRAADLLKHEALNPPREDQPRCTSLD SALLERKRLLSRKELELPENIADSSCTGSTEESEMLKRQRSLYIDLGALAGYFNLVRGPPTLEYG		

	SEQ ID NO: 229	1430 bp	
NOV52b, CG93659-03 DNA Sequence	CTGACACTGCACTGAGCACTTTATGAGCTTGAACTCTGTTAATCCTCACGACCACCTCATGAGACTC TCCAGAAAGAGCAACAGTAATGGAGTACATGAGCACTGGAAGTGACAATAAAGAAGAGATTGATTTA TTAATTAAACATTTAAATGTGTCTGATGTAATAGACATTATGGAAAATCTTTATGCAAGTGAAGAGC CAGCAGTTTATGAACCCAGTCTAATGACCATGTGTCAAGACAGTAATCAAAACGATGAGCGTTCTAA GTCTCTGCTGCTTAGTGGCCAAGAGGTACCATGGTTGTCATCAGTCAGATACGGAACTGTGGAGGAT TTGCTTGCTTTTGCAAACCATATATCCAACACTGCAAAGCATTTTTATGGACAACGACCACAGGAAT CTGGAATTTTATTAAACATGGTCATCACTCCCCAAAATGGACGTTACCAAATAGATTCCGATGTTCT CCTGATCCCCTGGAAGCTGACTTACAGGAATATTGGTTCTGATTTTATTTCTCGGGGCGCCTTTGGA AAGGTATACTTGGCACAAGATATAAAGACGAAGAAAAGAATGGCGTGTAAACTGATCCCAGTAGATC AATTTAAGCCATCTGATGTGGAAATCCAGGCTTGCTTCCGGCACGAGAACATCGCAGAGCTGTATGG CGCAGTCCTGTGGGGTGAAACTGTCCATCTCTTTATGGAAGCAGGCGAGGGAGGGTCTGTTCTGGAG AAACTGGAGAGCTGTGGACCAATGAGAGAATTTGAAATTATTTGGGTGACAAAGCATGTTCTCAAGG GACTTGATTTTCTACACTCAAAGAAAGTGATCCATCATGATATAAACATTTACATGAGCCCAGAGGT CATCCTGTGCAGGGGCCATTCAACCAAAGCAGACATCTACAGCCTGGGGGCCACGCTCATCCACATG CAGACGGGCACCCCACCCTGGGTGAAGCGCTACCCTCGCTCAGCCTATCCCTCCTACCTGTACATAA TCCACAAGCAAGCACCTCCACTGGAAGACATTGCAGATGACTGCAGTCCAGGGATGAGAGAGCTGAT AGAAGCTTCCCTGGAGAGAAACCCCAATCACCGCCCAAGAGCCGCAGACCTACTAAAACATGAGGCC CTGAACCCGCCCAGAGAGGATCAGCCACGCTGTCAGAGTCTGGACTCTGCCCTCTTGGAGCGCAAGA GGCTGCTGAGTAGGAAGGAGCTGGAACTTCCTGAGAACATTGCTGATTCTTCGTGCACAGGAAGCAC CGAGGAATCTGAGATGCTCAAGAGGCAACGCTCTCTCTACATCGACCTCGGCGCTCTGGCTGGCTAC TTCAATCTTGTTCGGGGACCACCAACGCTTGAATATGGCTGAAGGATGCCATGTTTGCTCTAAATTA AGACAGCATTGATCTCCTGGAGG		
	ORF Start: ATG at 87		ORF Stop: TGA at 1380

5

	SEQ ID NO: 230	431 aa	MW at 48882.2kD
NOV52b, CG93659-03 Protein Sequence	MEYMSTGSDNKEEIDLLIKHLNVSDVIDIMENLYASEEPAVYEPSLMTMCQDSNQNDERSKSLLLSG QEVPWLSSVRYGTVEDLLAFANHISNTAKHFYGQRPQESGILLNMVITPQNGRYQIDSDVLLIPWKL TYRNIGSDFISRGAPGKVYLAQDIKTKKRMACKLIPVDQFKPSDVEIQACFRHENIAELYGAVLWGE TVHLFMEAGEGGSVLEKLESCGPMREFEIIWVTKHVLKGLDFLHSKKVIHHDINIYMSPEVILCRGH STKADIYSLGATLIHMQTGTPPWVKRYPRSAYPSYLYIIHKQAPPLEDIADDCSPGMRELIEASLER NPNHRPRAADLLKHEALNPPREDQPRCQSLDSALLERKRLLSRKELELPENIADSSCTGSTEESEML KRQRSLYIDLGALAGYFNLVRGPPTLEYG		

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	SEQ ID NO: 231	1538 bp	
NOV52c, CG93659-02 DNA Sequence	CTGACACTGCACTGAGCACTTTATGAGCTTGAACTCTGTTAATCCTCACGACCACCTCATGAGACTC TCCAGAAAGAGCAACAGTAATGGAGTACATGAGCACTGGAAGTGACAATAAAGAAGAGATTGATTTA TTAATTAAACATTTAAATGTGTCTGATGTAATAGACATTATGGAAAATCTTTATGCAAGTGAAGAGC CAGCAGTTTATGAACCCAGTCTAATGACCATGTGTCAAGACAGTAATCAAAACGATGAGCGTTCTAA GTCTCTGCTGCTTAGTGGCCAAGAGGTACCATGGTTGTCATCAGTCAGATACGGAACTGTGGAGGAT TTGCTTGCTTTTGCAAACCATATATCCAACACTGCAAAGCATTTTTATGGACAACGACCACAGGAAT CTGGAATTTTATTAAACATGGTCATCACTCCCCAAAATGGACGTTACCAAATAGATTCCGATGTTCT CCTGATCCCCTGGAAGCTGACTTACAGGAATATTGGTTCTGATTTTATTTCTCGGGGCGCCTTTGGA AAGGTATACTTGGCACAAGATATAAAGACGAAGAAAAGAATGGCGTGTAAACTGATCCCAGTAGATC AATTTAAGCCATCTGATGTGGAAATCCAGGCTTGCTTCCGGCACGAGAACATCGCAGAGCTGTATGG CGCAGTCCTGTGGGGTGAAACTGTCCATCTCTTTATGGAAGCAGGCGAGGGAGGGTCTGTTCTGGAG AAACTGGAGAGCTGTGGACCAATGAGAGAATTTGAAATTATTTGGGTGACAAAGCATGTTCTCAAGG GACTTGATTTTCTACACTCAAAGAAAGTGATCCACCATGATATTAAACCTAGCAACATTGTTTTCAT GTCCACAAAAGCTGTTTTGGTGGATTTTGGCCTAAGTGTTCAAATGACCGAAGATGTCTATTTTCCT AAGGACCTCCGAGGAACAGAGATTTACATGAGCCCAGAGGTCATCCTGTGCAGTGGCCATTCAACCA AAGCAGACATCTACAGCCTGGGGGCCACGCTCATCCACATGCAGACGGGCACCCCACCCTGGGTGAA GCGCTACCCTCGCTCAGCCTATCCCTCCTACCTGTACATAATCCACAAGCAAGCACCTCCACTGGAA GACATTGCAGATGACTGCAGTCCAGGGATGAGAGAGCTGATAGAAGCTTCCCTGGAGAGAAACCCCA ATCACCGCCCAAGAGCCGCAGACCTACTAAAACATGAGGCCCTGAACCCGCCCAGAGAGGATCAGCC ACGCTGTCAGAGTCTGGACTCTGCCCTCTTGGAGCGCAAGAGGCTGCTGAGTAGGAAGGAGCTGGAA CTTCCTGAGAACATTGCTGATTCTTCGTGCACAGGAAGCACCGAGGAATCTGAGATGCTCAAGAGGC AACGCTCTCTCTACATCGACCTCGGCGCTCTGGCTGGCTACTTCAATCTTGTTCGGGGACCACCAAC		

	GCTTGAATATGGCTGAAGGATGCCATGTTTGCTCTAAATTAAGACAGCATTGATCTCCTGGAGG		
	ORF Start: ATG at 87		ORF Stop: TGA at 1488

	SEQ ID NO: 232	467 aa	MW at 52844.7kD
NOV52c, CG93659-02 Protein Sequence	MEYMSTGSDNKEEIDLLIKHLNVSDVIDIMENLYASEEPAVYEPSLMTMCQDSNQNDERSKSLLLSG QEVPWLSSVRYGTVEDLLAFANHISNTAKHFYGQRPQESGILLNMVITPQNGRYQIDSDVLLIPWKL TYRNIGSDFISRGAFGKVYLAQDIKTKKRMACKLIPVDQFKPSDVEIQACFRHENIAELYGAVLWGE TVHLFMEAGEGGSVLEKLESCGPMREFEIIWVTKHVLKGLDFLHSKKVIHHDIKPSNIVFMSTKAVL VDFGLSVQMTEDVYFPKDLRGTEIYMSPEVILCSGHSTKADIYSLGATLIHMQTGTPPWVKRYPRSA YPSYLYIIHKQAPPLEDIADDCSPGMRELIEASLERNPNHRPRAADLLKHEALNPPREDQPRCQSLD SALLERKRLLSRKELELPENIADSSCTGSTEESEMLKRQRSLYIDLGALAGYFNLVRGPPTLEYG

Sequence comparison of the above protein sequences yields the following sequence relationships shown in Table 52B.

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Table 52B. Comparison of NOV52a against NOV52b and NOV52c.		
Protein Sequence	NOV52a Residues/ Match Residues	Identities/ Similarities for the Matched Region
NOV52b	1..467 1..431	413/467 (88%) 413/467 (88%)
NOV52c	1..467 1..467	449/467 (96%) 449/467 (96%)

Further analysis of the NOV52a protein yielded the following properties shown in

15 Table 52C.

Table 52C. Protein Sequence Properties NOV52a	
PSort analysis:	0.6500 probability located in cytoplasm; 0.1000 probability located in mitochondrial matrix space; 0.1000 probability located in lysosome (lumen); 0.0000 probability located in endoplasmic reticulum (membrane)
SignalP analysis:	No Known Signal Sequence Predicted

20   A search of the NOV52a protein against the Geneseq database, a proprietary database that contains sequences published in patents and patent publication, yielded several homologous proteins shown in Table 52D.

Table 52D. Geneseq Results for NOV52a				
Geneseq Identifier	Protein/Organism/Length [Patent #, Date]	NOV52a Residues/ Match Residues	Identities/ Similarities for the Matched Region	Expect Value
AAE05951	Human cot oncoprotein encoded by D14497 oncogene - Homo sapiens, 467 aa. [US6265216-B1, 24-JUL-2001]	1..467 1..467	467/467 (100%) 467/467 (100%)	0.0
AAY79244	Human COT - Homo sapiens, 467 aa. [WO200011191-A2, 02-MAR-2000]	1..467 1..467	467/467 (100%) 467/467 (100%)	0.0
AAE10313	Human Tp12 protein - Homo sapiens, 467 aa. [WO200166559-A1, 13-SEP-2001]	1..467 1..467	466/467 (99%) 466/467 (99%)	0.0
AAE10314	Rat Tp12 protein - Rattus sp, 467 aa. [WO200166559-A1, 13-SEP-2001]	1..467 1..467	439/467 (94%) 454/467 (97%)	0.0
AAY79243	Rat TPL-2 - Rattus norvegicus, 467 aa. [WO200011191-A2, 02-MAR-2000]	1..467 1..467	438/467 (93%) 453/467 (96%)	0.0

5   In a BLAST search of public sequence datbases, the NOV52a protein was found to have homology to the proteins shown in the BLASTP data in Table 52E.

Table 52E. Public BLASTP Results for NOV52a				
Protein Accession Number	Protein/Organism/Length	NOV52a Residues/ Match Residues	Identities/ Similarities for the Matched Portion	Expect Value
P41279	Mitogen-activated protein kinase kinase kinase 8 (EC 2.7.1.-) (COT proto-oncogene serine/threonine-protein kinase) (C-COT) (Cancer Osaka thyroid oncogene) - Homo sapiens (Human), 467 aa.	1..467 1..467	467/467 (100%) 467/467 (100%)	0.0

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A48713	serine/threonine-specific protein kinase cot, 58K form - human, 467 aa.	1..467 1..467	466/467 (99%) 466/467 (99%)	0.0
Q63562	Mitogen-activated protein kinase kinase kinase 8 (EC 2.7.1.-) (Tumor progression locus 2) (TPL-2) - Rattus norvegicus (Rat), 467 aa.	1..467 1..467	438/467 (93%) 453/467 (96%)	0.0
Q07174	Mitogen-activated protein kinase kinase kinase 8 (EC 2.7.1.-) (COT proto-oncogene serine/threonine-protein kinase) (C-COT) (Cancer Osaka thyroid oncogene) - Mus musculus (Mouse), 467 aa.	1..467 1..467	435/467 (93%) 454/467 (97%)	0.0
A41253	kinase-related transforming protein (EC 2.7.1.-) - human, 415 aa.	1..397 1..397	379/397 (95%) 379/397 (95%)	0.0

PFam analysis predicts that the NOV52a protein contains the domains shown in the Table 52F.

Table 52F. Domain Analysis of NOV52a			
Pfam Domain	NOV52a Match Region	Identities/ Similarities for the Matched Region	Expect Value
pkinase	146..388	74/279 (27%) 187/279 (67%)	4.7e-54

Example 53.

The NOV53 clone was analyzed, and the nucleotide and encoded polypeptide sequences are shown in Table 53A.

Table 53A. NOV53 Sequence Analysis			
	SEQ ID NO: 233	1078 bp	
NOV53a, CG94521-01 DNA Sequence	GCGGCTACATTCGGCCCGGCCATGGCAGCGGCGCCCCTGAAAGTGTGCATCGTGGGCTCGGGGAACT GGGGTTCAGCTGTTGCAAAAATAATTGGTAATAACGTCAAGAAACTTCAGAAATTTGCCTCCACAGT CAAGATGTGGGTCTTTGAAGAAACAGTGAATGGCAGAAAACTGACAGACATCATAAATAATGACCAT GAAAATGTAAAATATCTTCCTGGACACAAGCTGCCAGAAAATGTGGTTGCCATGTCAAATCTTAGCG AGGCTGTGCAGGATGCAGACCTGCTGGTGTTTGTCATTCCCCACCAGTTCATTCACAGAATCTGTGA TGAGATCACTGGGAGAGTGCCCAAGAAAGCGCTGGGAATCACCCTCATCAAGGGCATAGACGAGGGC	

	CCCGAGGGGCTGAAACTCATTTCTGACATCATCCGTGAGAAGATGGGTATTGACATCAGTGTGCTGA TGGGAGCCAACATTGCCAATGAGGTGGCTGCAGAGAAGTTCTGTGAGACCACCATCGGCAGCAAAGT AATGGAGAACGGCCTTCTCTTCAAAGAACTTCTGCAGACTCCAAATTTTCGAATTACGGTGGTTGAT GATGCAGACACTGTTGAACTCTGTGGTGCGCTTAAGAACATCGTAGCTGTGGGAGCTGGGTTCTGCG ACGGCCTCCGCTGTGGAGACAACACCAAAGCGGCCGTCATCCGCCTGGGACTCATGGAAATGATTGC TTTTGCCAGGATCTTCTGCAAAGGCCAAGTGTCTACAGCCACCTTCCTAGAGAGCTGCGGGGTGGCC GACCTGATCACCACCTGTTACGGAGGGCGGAACCGCAGGGTGGCCGAGGCCTTCGCCAGAACTGGGA AGACCATTGAAGAGTTGGAGAAGGAGATGCTGAATGGGCAAAAGCTCCAAGGACCGCAGACTTCTGC TGAAGTGTACCGCATCCTCAAACAGAAGGGACTACTGGACAAGTTTCCATTGTTTACTGCAGTGTAT CAGATCTGCTACGAAAGCAGACCAGTTCAAGAGATGTTGTCTTGTCTTCAGAGCCATCCAGAGCATA CATAAA		
	ORF Start: ATG at 22		ORF Stop: TAA at 1075

	SEQ ID NO: 234	351 aa	MW at 38418.3kD
NOV53a, CG94521-01 Protein Sequence	MAAAPLKVCIVGSGNWGSAVAKIIGNNVKKLQKFASTVKMWVFEETVNGRKLTDIINNDHENVKYLP GHKLPENVVAMSNLSEAVQDADLLVFVIPHQFIHRICDEITGRVPKKALGITLIKGIDEGPEGLKLI SDIIREKMGIDISVLMGANIANEVAAEKFCETTIGSKVMENGLLFKELLQTPNFRITVVDDADTVEL CGALKNIVAVGAGFCDGLRCGDNTKAAVIRLGLMEMIAFARIFCKGQVSTATFLESCGVADLITTCY GGRNRRVAEAFARTGKTIEELEKEMLNGQKLQGPQTSAEVYRILKQKGLLDKFPLFTAVYQICYESR PVQEMLSCLQSHPEHT		

	SEQ ID NO: 235	936 bp	
NOV53b, CG94521-03 DNA Sequence	TACATTCGGCCCGGCCATGGCAGCGGCGCCCCTGAAAGTGTGCATCGTGGGCTCGGGGAACTGGGGT TCAGCTGTTGCAAAAATAATTGGTAATAATGTCAAGAAACTTCAGAAATTTGCCTCCACAGTCAAGA TGTGGGTCTTTGAAGAAACAGTGAATGGCAGAAAACTGACAGACATCATAAATAATGACCATGAAAA TGTAAAATATCTTCCTGGACACAAGCTGCCAGAAAATGTGGGCATAGACGAGGGCCCCGAGGGGCTG AAGCTCATTTCTGACATCATCCGTGAGAAGATGGGTATTGACATCAGTGTGCTGATGGGAGCCAACA CCTTCTCTTCAAAGAACTTCTGCAGACTCCAAATTTTCGAATTACCGTGGTTGATGATGCAGACACT GTTGAACTCTGTGGTGCGCTTAAGAACATCGTAGCTGTGGGAGCTGGGTTCTGCGACGGCCTCCGCT CTTCTGCAAAGGCCAAGTGTCTACAGCCACCTTCCTAGAGAGCTGCGGGGTGGCCGACCTGATCACC ACCTGTTACGGAGGGCGGAACCGCAGGGTGGCCGAGGCCTTCGCCAGAACTGGGAAGACCATTGAAG AGTTGGAGAAGGAGATGCTGAATGGGCAAAAGCTCCAAGGACCGCAGACTTCTGCTGAAGTGTACCG CATCCTCAAACAGAAGGGACTACTGGACAAGTTTCCATTGTTTACTGCAGTGTATCAGATCTGCTAC GAAAGCAGACCAGTTCAAGAGATGTTGTCTTGTCTTCAGAGCCATCCAGAGCATACATAAAAAGG		
	ORF Start: ATG at 17		ORF Stop: TAA at 929

	SEQ ID NO: 236	304 aa	MW at 33235.2kD
NOV53b, CG94521-03 Protein Sequence	MAAAPLKVCIVGSGNWGSAVAKIIGNNVKKLQKFASTVKMWVFEETVNGRKLTDIINNDHENVKYLP GHKLPENVGIDEGPEGLKLISDIIREKMGIDISVLMGANIANEVAAEKFCETTIGSKVMENGLLFKE LLQTPNFRITVVDDADTVELCGALKNIVAVGAGFCDGLRCGDNTKAAVIRLGLMEMIAFARIFCKGQ VSTATFLESCGVADLITTCYGGRNRRVAEAFARTGKTIEELEKEMLNGQKLQGPQTSAEVYRILKQK GLLDKFPLFTAVYQICYESRPVQEMLSCLQSHPEHT		

	SEQ ID NO: 237	1077 bp	

| NOV53c, CG94521-02 DNA Sequence | <u>TACATTCGGCCCCGGCC</u>**ATG**GCAGCGGCGCCCCTGAAAGTGTGCATCGTGGGCTCGGGGAACTGGGGT TCAGCTGTTGCAAAAATAATTGGTAATAATGTCAAGAAACTTCAGAAATTTGCCTCCACAGTCAAGA TGTGGGTCTTTGAAGAAACAGTGAATGGCAGAAAACTGACAGACATCATAAATAATGACCATGAAAA TGTAAAATATCTTCCTGGACACAAGCTGCCAGAAAATGTGGTTGCCATGTCAAATCTTAGCGAGGCT GTGCAGGATGCAGACCTGCTGGTGTTTGTCATTCCCCACCAGTTCATTCACAGAATCTGTGATGAGA TCACTGGGAGAGTGCCCAAGAAAGCGCTGGGAATCACCCTCATCAAGGGCATAGACGAGGGCCCCGA GGGGCTGAAGCTCATTTCTGACATCATCCGTGAGAAGATGGGTATTGACATCAGTGTGCTGATGGGA GCCAACATTGCCAATGAGGTGGCTGCAGAGAAGTTCTGTGAGACCACCATCGGCAGCAAAGTAATGG AGAACGGCCTTCTCTTCAAAGAACTTCTGCAGACTCCAAATTTTCGAATTACCGTGGTTGATGATGC AGACACTGTTGAACTCTGTGGTGCGCTTAAGAACATCGTAGCTGTGGGAGCTGGGTTCTGCGACGGC CTCCGCTGTGGAGACAACACCAAAGCGGCCGTCATCCGCCTGGGACTCATGGAAATGATTGCTTTTG CCAGGATCTTCTGCAAAGGCCAAGTGTCTACAGCCACCTTCCTAGAGAGCTGCGGGGTGGCCGACCT GATCACCACCTGTTACGGAGGGCGGAACCGCAGGGTGGCCGAGGCCTTCGCCAGAACTGGGAAGACC ATTGAAGAGTTGGAGAAGGAGATGCTGAATGGGCAAAAGCTCCAAGGACCGCAGACTTCTGCTGAAG TGTACCGCATCCTCAAACAGAAGGGACTACTGGACAAGTTTCCATTGTTTACTGCAGTGTATCAGAT CTGCTACGAAAGCAGACCAGTTCAAGAGATGTTGTCTTGTCTTCAGAGCCATCCAGAGCATACA**TAA** <u>AAAGG</u> |

	ORF Start: ATG at 17			ORF Stop: TAA at 1070

	SEQ ID NO: 238	351 aa	MW at 38418.3kD
NOV53c, CG94521-02 Protein Sequence	MAAAPLKVCIVGSGNWGSAVAKIIGNNVKKLQKFASTVKMWVFEETVNGRKLTDIINNDHENVKYLP GHKLPENVVAMSNLSEAVQDADLLVFVIPHQFIHRICDEITGRVPKKALGITLIKGIDEGPEGLKLI SDIIREKMGIDISVLMGANIANEVAAEKFCETTIGSKVMENGLLFKELLQTPNFRITVVDDADTVEL CGALKNIVAVGAGFCDGLRCGDNTKAAVIRLGLMEMIAFARIFCKGQVSTATFLESCGVADLITTCY GGRNRRVAEAFARTGKTIEELEKEMLNGQKLQGPQTSAEVYRILKQKGLLDKFPLFTAVYQICYESR PVQEMLSCLQSHPEHT		

Sequence comparison of the above protein sequences yields the following sequence relationships shown in Table 53B.

10

Table 53B. Comparison of NOV53a against NOV53b and NOV53c.		
Protein Sequence	NOV53a Residues/ Match Residues	Identities/ Similarities for the Matched Region
NOV53b	1..351 1..304	304/351 (86%) 304/351 (86%)
NOV53c	1..351 1..351	351/351 (100%) 351/351 (100%)

Further analysis of the NOV53a protein yielded the following properties shown in
15　Table 53C.

Table 53C. Protein Sequence Properties NOV53a

PSort analysis:	0.6500 probability located in cytoplasm; 0.1000 probability located in mitochondrial matrix space; 0.1000 probability located in lysosome (lumen); 0.0000 probability located in endoplasmic reticulum (membrane)
SignalP analysis:	Cleavage site between residues 22 and 23

A search of the NOV53a protein against the Geneseq database, a proprietary database that contains sequences published in patents and patent publication, yielded several homologous proteins shown in Table 53D.

Table 53D. Geneseq Results for NOV53a				
Geneseq Identifier	Protein/Organism/Length [Patent #, Date]	NOV53a Residues/ Match Residues	Identities/ Similarities for the Matched Region	Expect Value
ABB64184	Drosophila melanogaster polypeptide SEQ ID NO 19344 - Drosophila melanogaster, 360 aa. [WO200171042-A2, 27-SEP-2001]	3..350 2..349	212/349 (60%) 263/349 (74%)	e-120
AAG08446	Arabidopsis thaliana protein fragment SEQ ID NO: 5988 - Arabidopsis thaliana, 366 aa. [EP1033405-A2, 06-SEP-2000]	7..331 22..349	180/329 (54%) 233/329 (70%)	8e-95
AAG08445	Arabidopsis thaliana protein fragment SEQ ID NO: 5987 - Arabidopsis thaliana, 400 aa. [EP1033405-A2, 06-SEP-2000]	7..331 56..383	180/329 (54%) 233/329 (70%)	8e-95
AAG08444	Arabidopsis thaliana protein fragment SEQ ID NO: 5986 - Arabidopsis thaliana, 421 aa. [EP1033405-A2, 06-SEP-2000]	7..331 77..404	180/329 (54%) 233/329 (70%)	8e-95
AAG39422	Arabidopsis thaliana protein fragment SEQ ID NO: 48774 - Arabidopsis thaliana, 366 aa. [EP1033405-A2, 06-SEP-2000]	7..331 22..349	180/329 (54%) 232/329 (69%)	1e-94

In a BLAST search of public sequence datbases, the NOV53a protein was found to
have homology to the proteins shown in the BLASTP data in Table 53E.

Table 53E. Public BLASTP Results for NOV53a				
Protein Accession Number	Protein/Organism/Length	NOV53a Residues/ Match Residues	Identities/ Similarities for the Matched Portion	Expect Value
AAH28726	KIAA0089 protein - Homo sapiens (Human), 351 aa.	1..351 1..351	351/351 (100%) 351/351 (100%)	0.0
Q14702	KIAA0089 protein - Homo sapiens (Human), 411 aa (fragment).	1..351 61..411	351/351 (100%) 351/351 (100%)	0.0
O57656	Glycerol-3-phosphate dehydrogenase [NAD+], cytoplasmic (EC 1.1.1.8) (GPD-C) (GPDH-C) - Fugu rubripes (Japanese pufferfish) (Takifugu rubripes), 351 aa.	3..350 2..350	265/349 (75%) 306/349 (86%)	e-155
Q98SJ9	Glycerol-3-phosphate dehydrogenase (EC 1.1.1.8) - Salmo salar (Atlantic salmon), 350 aa.	7..350 5..349	258/345 (74%) 301/345 (86%)	e-152
AAH32234	Glycerol-3-phosphate dehydrogenase 1 (soluble) - Homo sapiens (Human), 349 aa.	4..350 2..348	249/347 (71%) 297/347 (84%)	e-149

5

PFam analysis predicts that the NOV53a protein contains the domains shown in the
Table 53F.

10

Table 53F. Domain Analysis of NOV53a			
Pfam Domain	NOV53a Match Region	Identities/ Similarities for the Matched Region	Expect Value
NAD_Gly3P_dh	5..344	167/365 (46%) 307/365 (84%)	2.1e-184

Example 54.

The NOV54 clone was analyzed, and the nucleotide and encoded polypeptide
sequences are shown in Table 54A.

5

Table 54A. NOV54 Sequence Analysis			
	SEQ ID NO: 239	1552 bp	
NOV54a, CG96613-01 DNA Sequence	TTATTCCCCACTTTACCTGGCTAATTGAAGTGTAACAAAAGCTTCATCCAGGAACATTGGCGCGGGA AACCTGGCGTACTGGCTGTGGCTTCTCTAGCGGGACTCGGC**ATG**AGGCTGGCGCGGCTGCTTCGCGG AGCCGCCTTGGCCGGCCCGGGCCCGGGGCTGCGCGCCGCCGGCTTCAGCCGCAGCTTCAGCTCGGAC TCGGGCTCCAGCCCGGCGTCCGAGCGCGGCGTTCCGGGCCAGGTGGACTTCTACGCGCGCTTCTCGC CGTCCCCGCTCTCCATGAAGCAGTTCCTGGACTTCGGATCAGTGAATGCTTGTGAAAAGACCTCATT TATGTTTCTGCGGCAAGAGTTGCCTGTCAGACTGGCAAATATAATGAAAGAAATAAGTCTCCTTCCA GATAATCTTCTCAGGACACCATCCGTTCAATTGGTACAAAGCTGGTATATCCAGAGTCTTCAGGAGC TTCTTGATTTTAAGGACAAAAGTGCTGAGGATGCTAAAGCTATTTATGACTTTACAGATACTGTGAT ACGGATCAGAAACCGACACAATGATGTCATTCCCACAATGGCCCAGGGTGTGATTGAATACAAGGAG AGCTTTGGGGTGGATCCTGTCACCAGCCAGAATGTTCAGTACTTTTTGGATCGATTCTACATGAGTC GCATTTCAATTAGAATGTTACTCAATCAGCACTCTTTATTGTTTGGTGGAAAAGGCAAAGGAAGTCC ATCTCATCGAAAACACATTGGAAGCATAAATCCAAACTGCAATGTACTTGAAGTTATTAAAGATGGC TATGAAAATGCTAGGCGTCTGTGTGATTTGTATTATATTAACTCTCCCGAACTAGAACTTGAAGAAC TAAATGCAAAATCACCAGGACAGCCAATCAAGTGGTTTATGTACCATCCCATCTCTATCACATGGT GTTTGAACTTTTCAAGAATGCAATGAGAGCCACTATGGAACACCATGCCAACAGAGGTGTTTACCCC CCTATTCAAGTTCATGTCACGCTGGGTAATGAGGATTTGACTGTGAAGATGAGTGACCGAGGAGGTG GCGTTCCTTTGAGGAAAATTGACAGACTTTTCAACTACATGTATTCAACTGCACCAAGACCTCGTGT TGAGACCTCCCGCCAGTGCCTCTGGCTGGTTTTGGTTTATGGATTGCCCATATCACGTCTTTACGCA CAATACTTCCAAGGAGACCTGAAGCTGTATTCCCTAGAGGGTTACGGGACAGATGCAGTTATCTACA TTAAGGCTCTGTCAACAGACTCAATAGAAAGACTCCCAGTGTATAACAAAGCTGCCTGGAAGCATTA CAACACCAACCACGAGGCTGATGACTGGTGCGTCCCCAGCAGAGAACCCAAAGACATGACGACGTTC CGCAGTGCC**TAG**ACACACTGGGGACATCGGAAAATCCAAATGTGGCTTTTGTATTAAATTTGGAAGG TATGGTGTTCAGAACTATATTATACCAAGTACTTTATTTATCGTTTTCACAAAACTATTTGAGTAGA ATAAATGGAAA		
	ORF Start: ATG at 109		ORF Stop: TAG at 1417

	SEQ ID NO: 240	436 aa	MW at 49243.6kD
NOV54a, CG96613-01 Protein Sequence	MRLARLLRGAALAGPGPGLRAAGFSRSFSSDSGSSPASERGVPGQVDFYARFSPSPLSMKQFLDFGS VNACEKTSFMFLRQELPVRLANIMKEISLLPDNLLRTPSVQLVQSWYIQSLQELLDFKDKSAEDAKA IYDFTDTVIRIRNRHNDVIPTMAQGVIEYKESFGVDPVTSQNVQYFLDRFYMSRISIRMLLNQHSLL FGGKGKGSPSHRKHIGSINPNCNVLEVIKDGYENARRLCDLYYINSPELELEELNAKSPGQPIQVVY VPSHLYHMVFELFKNAMRATMEHHANRGVYPPIQVHVTLGNEDLTVKMSDRGGGVPLRKIDRLFNYM YSTAPRPRVETSRAVPLAGFGYGLPISRLYAQYFQGDLKLYSLEGYGTDAVIYIKALSTDSIERLPV YNKAAWKHYNTNHEADDWCVPSREPKDMTTFRSA		

	SEQ ID NO: 241	1612 bp	
NOV54b, CG96613-03 DNA Sequence	TTATTCCCCACTTTACCTGGCTAATTGAAGTGTAACAAAAGCTTCATCCAGGAACATTGGCGCGGGA AACCTGGCGTACTGGCTGTGGCTTCTCTAGCGGGACTCGGC**ATG**AGGCTGGCGCGGCTGCTTCGCGG AGCCGCCTTGGCCGGCCCGGGCCCGGGGCTGCGCGCCGCCGGCTTCAGCCGCAGCTTCAGCTCGGAC TCGGGCTCCAGCCCGGCGTCCGAGCGCGGCGTTCCGGGCCAGGTGGACTTCTACGCGCGCTTCTCGC CGTCCCCGCTCTCCATGAAGCAGTTCCTGGACTTCGGATCAGTGAATGCTTGTGAAAAGACCTCATT TATGTTTCTGCGGCAAGAGTTGCCTGTCAGACTGGCAAATATAATGAAAGAAATAAGTCTCCTTCCA GATAATCTTCTCAGGACACCATCCGTTCAATTGGTACAAAGCTGGTATATCCAGAGTCTTCAGGAGC TTCTTGATTTTAAGGACAAAAGTGCTGAGGATGCTAAAGCTATTTATGAAAGGCCTAGAAGAACATG GTTGCAGGTCTCTAGTTTATGCTGTATGGCCTGCAAGATGATCTTTACAGATACTGTGATACGGATC AGAAACCGACACAATGATGTCATTCCCACAATGGCCCAGGGTGTGATTGAATACAAGGAGAGCTTTG GGGTGGATCCTGTCACCAGCCAGAATGTTCAGTACTTTTTGGATCGATTCTACATGAGTCGCATTTC		

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	AATTAGAATGTTACTCAATCAGCACTCTTTATTGTTTGGTGGAAAAGGCAAAGGAAGTCCATCTCAT
	CGAAAACACATTGGAAGCATAAATCCAAACTGCAATGTACTTGAAGTTATTAAAGATGGCTATGAAA
	ATGCTAGGCGTCTGTGTGTGATTTGTATTATATTAACTCTCCCGAACTAGAACTTGAAGAACTAAATGC
	AAAATCACCAGGACAGCCAATACAAGTGGTTTATGTACCATCCCATCTCTATCACATGGTGTTTGAA
	CTTTTCAAGAATGCAATGAGAGCCACTATGGAACACCATGCCAACAGAGGTGTTTACCCCCCTATTC
	AAGTTCATGTCACGCTGGGTAATGAGGATTTGACTGTGAAGATGAGTGACCGAGGAGGTGGCGTTCC
	TTTGAGGAAAATTGACAGACTTTTCAACTACATGTATTCAACTGCACCAAGACCTCGTGTTGAGACC
	TCCCGCGCAGTGCCTCTGGCTGGTTTTGGTTATGGATTGCCCATATCACGTCTTTACGCACAATACT
	TCCAAGGAGACCTGAAGCTGTATTCCCTAGAGGGTTACGGGACAGATGCAGTTATCTACATTAAGGC
	TCTGTCAACAGACTCAATAGAAAGACTCCCAGTGTATAACAAAGCTGCCTGGAAGCATTACAACACC
	AACCACGAGGCTGATGACTGGTGCGTCCCCAGCAGAGAACCCAAAGACATGACGACGTTCCGCAGTG
	CCTAGACACACTGGGGACATCGGAAAATCCAAATGTGGCTTTTGTATTAAATTTGGAAGGTATGGTG
	TTCAGAACTATATTATACCAAGTACTTTATTTATCGTTTTCACAAAACTATTTGAGTAGAATAAATG
	GAAA

ORF Start: ATG at 109		ORF Stop: TAG at 1477

	SEQ ID NO: 242	456 aa	MW at 51622.6kD
NOV54b, CG96613-03 Protein Sequence	MRLARLLRGAALAGPGPGLRAAGFSRSFSSDSGSSPASERGVPGQVDFYARFSPSPLSMKQFLDFGS VNACEKTSFMFLRQELPVRLANIMKEISLLPDNLLRTPSVQLVQSWYIQSLQELLDFKDKSAEDAKA IYERPRRTWLQVSSLCCMACKMIFTDTVIRIRNRHNDVIPTMAQGVIEYKESFGVDPVTSQNVQYFL DRFYMSRISIRMLLNQHSLLFGGKGKGSPSHRKHIGSINPNCNVLEVIKDGYENARRLCDLYYINSP ELELEELNAKSPGQPIQVVYVPSHLYHMVFELFKNAMRATMEHHANRGVYPPIQVHVTLGNEDLTVK MSDRGGGVPLRKIDRLFNYMYSTAPRPRVETSRAVPLAGFGYGLPISRLYAQYFQGDLKLYSLEGYG TDAVIYIKALSTDSIERLPVYNKAAWKHYNTNHEADDWCVPSREPKDMTTFRSA		

	SEQ ID NO: 243	967 bp	
NOV54c, CG96613-02 DNA Sequence	TTATTCCCCACTTTACCTGGCTAATTGAAGTGTAACAAAAGCTTCATCCAGGACAATTGGCGCGGGA AACCTGGCGTACTGGCTGGTTCTCTAGCGGGACTCGGCATGAGGCTGGCGCGGCTGCTTCGCGG AGCCGCCTTGGCCGGCCCGGGCCCGGGGCTGCGCGCCGCCGGCTTCAGCCGCAGCTTCAGCTCGGAC TCGGGCTCCAGCCCGGCGTCCGAGCGCGGCGTTCCGGGCCAGGTGGACTTCTACGCGCGCTTCTCGC CGTCCCCGCTCTCCATGAAGCAGTTCCTGGACTTCGGATCAGTGAATGCTTGTGAAAAGACCTCATT TATGTTTCTGCGGCAAGAGTTGCCTGTCAGACTGGCAAATATAATGAAAGAAATAAGTCTCCTTCCA GATAATCTTCTCAGGACACCATCCGTTCAATTGGTACAAAGCTGGTATATCCAGAGTCTTCAGGAGC TTCTTGATTTTAAGGACAAAAGTGCTGAGGATGCTAAAGCTATTTATGAAAGGCCTAGAAGAACATG GTTGCAGGTCTCTAGTTTATGCTGTATGGCCTGCAAGATGATCTTTACAGATACTGTGATACGGATC AGAAACCGACACAATGATGTCATTCCCACAATGGCCCAGGGTGTGATTGAATACAAGGAGAGCTTTG GGGTGGATCCTGTCACCAGCCAGAATGTTCAGTACTTTATTTATCGTTTTCACAAAACTATTTGAGT AGAATAAATGGAAAACTGAATTCCATTTGTGCCCGTTAAACCTCCTAAAGGATGAAATTGCACCTATT TTACACCTATATTTTCACAGTTAATTGAACATATTTTTAAACAACTGTAGTTTTGGGCAACTTTTCA CTTTGTGGTAGACTTCAGAAGTGTGGAAATCTTCGGGTTTCTATAGGAAACTAGTTTTTTTTTTTTT AAAAAAATCCTTTCTTTTTTGTGGGCTAG		

ORF Start: ATG at 109		ORF Stop: TGA at 733

	SEQ ID NO: 244	208 aa	MW at 23483.8kD
NOV54c, CG96613-02 Protein Sequence	MRLARLLRGAALAGPGPGLRAAGFSRSFSSDSGSSPASERGVPGQVDFYARFSPSPLSMKQFLDFGS VNACEKTSFMFLRQELPVRLANIMKEISLLPDNLLRTPSVQLVQSWYIQSLQELLDFKDKSAEDAKA IYERPRRTWLQVSSLCCMACKMIFTDTVIRIRNRHNDVIPTMAQGVIEYKESFGVDPVTSQNVQYFI YRFHKTI		

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Sequence comparison of the above protein sequences yields the following sequence relationships shown in Table 54B.

Table 54B. Comparison of NOV54a against NOV54b and NOV54c.		
Protein Sequence	NOV54a Residues/ Match Residues	Identities/ Similarities for the Matched Region
NOV54b	42..436 42..456	394/415 (94%) 395/415 (94%)
NOV54c	42..185 42..205	140/164 (85%) 143/164 (86%)

Further analysis of the NOV54a protein yielded the following properties shown in Table 54C.

Table 54C. Protein Sequence Properties NOV54a	
PSort analysis:	0.4251 probability located in mitochondrial matrix space; 0.3802 probability located in microbody (peroxisome); 0.1914 probability located in lysosome (lumen); 0.1017 probability located in mitochondrial inner membrane
SignalP analysis:	Cleavage site between residues 22 and 23

A search of the NOV54a protein against the Geneseq database, a proprietary database that contains sequences published in patents and patent publication, yielded several homologous proteins shown in Table 54D.

Table 54D. Geneseq Results for NOV54a				
Geneseq Identifier	Protein/Organism/Length [Patent #, Date]	NOV54a Residues/ Match Residues	Identities/ Similarities for the Matched Region	Expect Value
ABG16621	Novel human diagnostic protein #16612 - Homo sapiens, 415 aa. [WO200175067-A2, 11-OCT-2001]	42..435 21..413	269/395 (68%) 331/395 (83%)	e-162

ABB58044	Drosophila melanogaster polypeptide SEQ ID NO 924 - Drosophila melanogaster, 413 aa. [WO200171042-A2, 27-SEP-2001]	26..420 2..396	219/401 (54%) 288/401 (71%)	e-121
AAE07838	Maize pyruvate dehydrogenase kinase (PDK)-2 - Zea mays, 364 aa. [US6265636-B1, 24-JUL-2001]	40..401 8..364	144/374 (38%) 211/374 (55%)	2e-60
AAW64724	A. thaliana PDHK protein from clone YA5 - Arabidopsis thaliana, 366 aa. [WO9835044-A1, 13-AUG-1998]	57..401 29..366	142/357 (39%) 209/357 (57%)	3e-58
AAE07837	Maize pyruvate dehydrogenase kinase (PDK)-1 - Zea mays, 347 aa. [US6265636-B1, 24-JUL-2001]	40..401 8..347	135/371 (36%) 205/371 (54%)	4e-56

In a BLAST search of public sequence datbases, the NOV54a protein was found to have homology to the proteins shown in the BLASTP data in Table 54E.

5

Table 54E. Public BLASTP Results for NOV54a				
Protein Accession Number	Protein/Organism/Length	NOV54a Residues/ Match Residues	Identities/ Similarities for the Matched Portion	Expect Value
Q15118	[Pyruvate dehydrogenase [lipoamide]] kinase isozyme 1, mitochondrial precursor (EC 2.7.1.99) (Pyruvate dehydrogenase kinase isoform 1) - Homo sapiens (Human), 436 aa.	1..436 1..436	436/436 (100%) 436/436 (100%)	0.0
Q63065	[Pyruvate dehydrogenase [lipoamide]] kinase isozyme 1, mitochondrial precursor (EC 2.7.1.99) (Pyruvate dehydrogenase kinase isoform 1) (PDK P48) - Rattus norvegicus (Rat), 434 aa.	1..436 1..434	402/436 (92%) 412/436 (94%)	0.0

Q8R2U8	Similar to pyruvate dehydrogenase kinase, isoenzyme 1 - Mus musculus (Mouse), 432 aa.	1..436 1..432	401/436 (91%) 412/436 (93%)	0.0
Q15119	[Pyruvate dehydrogenase [lipoamide]] kinase isozyme 2, mitochondrial precursor (EC 2.7.1.99) (Pyruvate dehydrogenase kinase isoform 2) - Homo sapiens (Human), 407 aa.	37..434 11..405	277/398 (69%) 340/398 (84%)	e-168
I70159	[pyruvate dehydrogenase (lipoamide)] kinase (EC 2.7.1.99) 2 - human, 407 aa.	37..434 11..405	276/398 (69%) 340/398 (85%)	e-168

PFam analysis predicts that the NOV54a protein contains the domains shown in the Table 54F.

5

Table 54F. Domain Analysis of NOV54a			
Pfam Domain	NOV54a Match Region	Identities/ Similarities for the Matched Region	Expect Value
HATPase_c	268..393	32/134 (24%) 84/134 (63%)	8.5e-20

Example 55.

10   The NOV55 clone was analyzed, and the nucleotide and encoded polypeptide sequences are shown in Table 55A.

Table 55A. NOV55 Sequence Analysis			
	SEQ ID NO: 245	2885 bp	
NOV55a, CG96736-01 DNA Sequence	CGGCACGCCCGGGAGGCTTTCTCTGGCTGGTAACCGCTACTCCCGGACACCAGACCACCGCCTTCCG TACACAGGGGCCCGCATCCCACCCTCCCGGACCTAAGAGCCTGGGTCCCCTGTTTCCGGAGTCCGCT TCCCGGCCCCCAGATTCTGGCATCCCAGCCCTCAGTGTCCAAGACCCAGGCAGCCCGGGTCCCCGCC TCCCGGATCCAGGCGTCCGGGATCTGCGCCACCAGAACCTAGCCTCCTGCAGACCTCCGCCATCTGG GGGCACTCAACCTCCTGGAGCCAAGGGCCCCACGTCCCACCCAGAGAAACTCTCGTATTCCCAGCTC CTAGGGCCAAGGAACCCGGGCGCTCCGAACTCCCAGCTTTCGGACATCTGGCACACGGGGCAGAGCA GAGAAGCCTCAGCGCCCAGCCTGGGGAATTTAAACACTCCAGCTTCCAAGAGCCAAGGAACTTCAGT GCTGTGAACTCACAACTCTAAGGAGCCCTCCAAAGTTCCAGTCTCCAGGTGCTGTTACTCAACTCAG TCCTAGGAACGTCGGGTCCTGGGAAGGAGCCCAAGCGCTCCCAGCCAGCTTCCAGGCGCTAAGAAAC CCCGGTGCTTCCCATCATGGTGGCCGATCCTCCTCGAGACTCCAAGGGGCTCGCAGCGGCGGAGCCA CCGCCAACGGGGGCCTGGCAGCTGGCCTCCATCGAGGACCAAGGCGCGGCAGCAGGCGGCTACTGCG GTTCCCGGGACCTGGTGCGCCGCTGCCTTCGAGCCAACCTGCTTGTGCTGCTGACAGTGGTGGCCGT

315

	GGTGGCCGGCGTGGCGCTGGGACTGGGGGTGTCGGGGGGCCGGGGGGTGCGCTGGCGTTGGGCCCGGGA GCGCTTGAGGCCTTCGTCTTCCCGGGCGAGCTGCTGCTGCGTCTGCTGCGGATGATCATCTTGCCGC TGGTGGTGTGCAGCTTGATCGGCGGCGCCGCCAGCCTGGACCCCGGCGCGCTCGGCCGTCTGGGCGC CTGGGCGCTGCTCTTTTTCCTGGTCACCACGCTGCTGGCGTCGGCGCTCGGAGTGGGCTTGGCGCTG GCTCTGCAGCCGGGCGCCGCCTCCGCCGCCATCAACGCCTCCGTGGGAGCCGCGGGCAGTGCCGAAA ATGCCCCCAGCAAGGAGGTGCTCGATTCGTTCCTGGATCTTGCGAGAAATATCTTCCCTTCCAACCT GGTGTCAGCAGCCTTTCGCTCATACTCTACCACCTATGAAGAGAGGAATATCACCGGAACCAGGGTG AAGGTGCCCGTGGGGCAGGAGGTGGAGGGGATGAACATCCTGGGCTTGGTAGTGTTTGCCATCGTCT TTGGTGTGGCGCTGCGGAAGCTGGGGCCTGAAGGGGAGCTGCTTATCCGCTTCTTCAACTCCTTCAA TGAGGCCACCATGGTTCTGGTCTCCTGGATCATGTGGTACGCCCCTGTGGGCATCATGTTCCTGGTG GCTGGCAAGATCGTGGAGATGGAGGATGTGGGTTTACTCTTTGCCCGCCTTGGCAAGTACATTCTGT GCTGCCTGCTGGGTCACGCCATCCATGGGCTCCTGGTACTGCCCCTCATCTACTTCCTCTTCACCCG CAAAAACCCCTACCGCTTCCTGTGGGGCATCGTGACGCCGCTGGCCACTGCCTTTGGGACCTCTTCC AGTTCCGCCACGCTGCCGCTGATGATGAAGTGCGTGGAGGAGAATAATGGCGTGGCCAAGCACATCA GCCGTTTCATCCTGCCCATCGGCGCCACCGTCAACATGGACGGTGCCGCCCTCTTCCAGTGCGTGGC CGCAGTGTTCATTGCACAGCTCAGCCAGCAGTCCTTGGACTTCGTAAAGATCATCACCATCCTGGTC ACGGCCACAGCCGTCCAGCGTGGGGGCAGCGGGCATCCCTGCTGGAGGTGTCCTCACTCTGGCCATCA TCCTCGAAGCAGTCAACCTCCCGGTCGACCATATCTCCTTGATCCTGGCTGTGGACTGGCTAGTCGA CCGGTCCTGTACCGTCCTCAATGTAGAAGGTGACGCTCTGGGGGCAGGACTCCTCCAAAATTATGTG GACCGTACGGAGTCGAGAAGCACAGAGCCTGAGTTGATACAAGTGAAGAGTGAGCTGCCCCTGGATC CGCTGCCAGTCCCCACTGAGGAAGGAAACCCCCTCCTCAAACACTATCGGGGGCCCGCAGGGGATGC CACGGTCGCCTCTGAGAAGGAATCAGTCATG**TAA**ACCCCGGGAGGGGACCTTCCCTGCCCTGCTGGGG GTGCTCTTTGGACACTGGATTATGAGGAATGGATAAATGGATGAGCTAGGGCTCTGGGGGTCTGCCT GCACACTCTGGGGAGCCAGGGGCCCCAGCACCCTCCAGGACAGGAGATCTGGGATGCCTGGCTGCTG GAGTACATGTGTTCACAAGGGTTACTCCTCAAAACCCCCAGTTCTCACTCATGTCCCCAACTCAAGG CTAGAAAAACAGCAAGATGGAGAAATAATGTTCTGCTGCGTCCCCACCGTGACCTGCCTGGCCTCCCC TGTCTCAGGGAGCAGGTCACAGGTCACCATGGGGAATTCTAGCCCCCACTGGGGGGGATGTTACAACA CCATGCTGGTTATTTTGGCGGCTGTAGTTGTGGGGGGATGTGTGTGTGCACGTGTGTGTGTGTGTGT GTGTGTGTGTGTGTGTGTTCTGTGGCCTCCTGTCCCCATGGTACGTCCCACCCTGTCCCCAGATC CCCTATTCCCTCCACAATAACAGAAACACTCCCAGGGACTCTGGGGGAGAGGCTGAGGACAAATACCT GCTGTCACTCCAGAGGACATTTTTTTTTAGCAATAAAATTGAGTGTCAACTATTAAAAAAAAAAAAAA AAAA
	ORF Start: ATG at 620		ORF Stop: TAA at 2243

	SEQ ID NO: 246	541 aa	MW at 56620.6kD
NOV55a, CG96736-01 Protein Sequence	MVADPPRDSKGLAAAEPPPTGAWQLASIEDQGAAAGGYCGSRDLVRRCLRANLLVLLTVVAVVAGVA LGLGVSGAGGALALGPGALEAFVFPGELLLRLLRMIILPLVVCSLIGGAASLDPGALGRLGAWALLF FLVTTLLASALGVGLALALQPGAASAAINASVGAAGSAENAPSKEVLDSFLDLARNIFPSNLVSAAF RSYSTTYEERNITGTRVKVPVGQEVEGMNILGLVVFAIVFGVALRKLGPEGELLIRFFNSFNEATMV LVSWIMWYAPVGIMFLVAGKIVEMEDVGLLFARLGKYILCCLLGHAIHGLLVLPLIYFLFTRKNPYR FLWGIVTPLATAFGTSSSSATLPLMMKCVEENNGVAKHISRFILPIGATVNMDGAALFQCVAAVFIA QLSQQSLDFVKIITILVTATASSVGAAGIPAGGVLTLAIILEAVNLPVDHISLILAVDWLVDRSCTV LNVEGDALGAGLLQNYVDRTESRSTEPELIQVKSELPLDPLPVPTEEGNPLLKHYRGPAGDATVASE KESVM		

	SEQ ID NO: 247	2017 bp	
NOV55b, CG96736-02 DNA Sequence	CGTACAACTCCGCCCCATTGACGCAAATGGGCGGTAGGCGTGTACGGTGGGAGGTCTATATAAGCAG AGCTCTCTGGCTAACTAGAGAACCCACTGCTTACTGGCTTATCGAAATTAATACGACTCACTATAGG GAGACCCAAGCTGGCTAGCGTTTAAACTTAAGCTTGGTACCGAGCTCGGATCCACTAGTCCAGTGTG GTGGAATTCCACCATGGTGGCCGATCCTCCTCGAGACTCCAAGGGGCTCGCAGCGGCGGAGCCCACC GCCAACGGGGGCCTGGCGCTGGCCTCCATCGAGGACCAAGGCGCGGCAGCAGGCGGCTACTGCGGTT CCCGGGACCAGGTGCGCCGCTGCCTTCGAGCCAACCTGCTTGTGCTGCTGACAGTGGTGGCCGTGGT GGCCGGCGTGGCGCTGGGACTGGGGGTGTCGGGGGCCGGGGGGTGCGCTGGCGTTGGGCCCGGAGCGC TTGAGCGCCTTCGTCTTCCCGGGCGAGCTGCTGCTGCGTCTGCTGCGGATGATCATCTTGCCGCTGG TGGTGTGCAGCTTGATCGGCGGCGCCGCCAGCCTGGACCCCGGCGCGCTCGGCCGTCTGGGCGCCTG GGCGCTGCTCTTTTTCCTGGTCACCACGCTGCTGGCGTCGGCGCTCGGAGTGGGCTTGGCGCTGGCT CTGCAGCCGGGCGCCGCCTCCGCCGCCATCAACGCCTCCGTGGGAGCCGCGGGCAGTGCCGAAAATG CCCCCAGCAAGGAGGTGCTCGATTCGTTCCTGGATCTTGCGAGAAATATCTTCCCTTCCAACCTGGT GTCAGCAGCCTTTCGCTCATACTCTACCACCTATGAAGAGAGGAATATCACCGGAACCAGGGTGAAG GTGCCCGTGGGGCAGGAGGTGGAGGGGATGAACATCCTGGGCTTGGTAGTGTTTGCCATCGTCTTTG GTGTGGCGCTGCGGAAGCTGGGGCCTGAAGGGGAGCTGCTTATCCGCTTCTTCAACTCCTTCAATGA

	GGCCACCATGGTTCTGGTCTCCTGGATCATGTGGTATGCCCCTGTGGGCATCATGTTCCTGCTGGCT
	GGCAAGATCGTGGAGATGGAGGATGTGGGTTTACTCTTTGCCCGCCTTGGCAAGTACATTCTGTGCT
	GCCTGCTGGGTCACGCCATCCATGGGCTCCTGGTACTGCCCCTCATCTACTTCCTCTTCACCCGCAA
	AAACCCCTACCGCTTCCTGTGGGGCATCGTGACGCCGCTGGCCACTGCCTTTGGGACCTCTTCCAGT
	TCCGCCACGCTGCCGCTGATGATGAAGTGCGTGGAGGAGAATAATGGCGTGGCCAAGCACATCAGCC
	GTTTCATCCTGCCCATCGGCGCCACCGTCAACATGGACGGTGCCGCGCTCTTCCAGTGCGTGGCCGC
	AGTGTTCATTGCACAGCTCAGCCAGCAGTCCTTGGACTTCGTAAAGATCATCACCATCCTGGTCACG
	GCCACAGCGTCCAGCGTGGGGGCAGCGGGCATCCCTGCTGGAGGTGTCCTCACTCTGGCCATCATCC
	TCGAAGCAGTCAACCTCCCGGTCGACCATATCTCCTTGATCCTGGCTGTGGACTGGCTAGTCGACCG
	GTCCTGTACCGTCCTCAATGTAGAAGGTGACGCTCTGGGGGCAGGACTCCTCCAAAATTACGTGGAC
	CGTACGGAGTCGAGAAGCACAGAGCCTGAGTTGATACAAGTGAAGAGTGAGCTGCCCCTGGATCCGC
	TGCCAGTCCCCACTGAGGAAGGAAACCCCCTCCTCAAACACTATCGGGGGCCCGCAGGGGATGCCAC
	GGTCGCCTCTGAGAAGGAATCAGTCATGTAAGCGGCCGCTCGAGTCTAGAGGGCCCGTTTAAACCCG
	CTGATCAGCCTCGACTGTGCCTTCTAGTTGCCAGCCATCTGTTGTTTGCCCCTCCCCCGTGCCTTCC
	TTGACCCTGGAAGGTGCCACTCCCACTGTCCTTTCCTAATAAAATGAGGAAATTGCATCGCATTGTC
	TGAGTAG
ORF Start: at 134			ORF Stop: TAA at 1838

	SEQ ID NO: 248	568 aa	MW at 59557.8kD
NOV55b, CG96736-02 Protein Sequence	GDPSWLAFKLKLGTELGSTSPVWWNSTMVADPPRDSKGLAAAEPTANGGLALASIEDQGAAAGGYCG SRDQVRRCLRANLLVLLTVVAVVAGVALGLGVSGAGGALALGPERLSAFVFPGELLLRLLRMIILPL VVCSLIGGAASLDPGALGRLGAWALLFFLVTTLLASALGVGLALALQPGAASAAINASVGAAGSAEN APSKEVLDSFLDLARNIFPSNLVSAAFRSYSTTYEERNITGTRVKVPVGQEVEGMNILGLVVFAIVF GVALRKLGPEGELLIRFFNSFNEATMVLVSWIMWYAPVGIMPLVAGKIVEMEDVGLLFARLGKYILC CLLGHAIHGLLVLPLIYFLFTRKNPYRFLWGIVTPLATAFGTSSSSATLPLMMKCVEENNGVAKHIS RFILPIGATVNMDGAALFQCVAAVFIAQLSQQSLDFVKIITILVTATASSVGAAGIPAGGVLTLAII LEAVNLPVDHISLILAVDWLVDRSCTVLNVEGDALGAGLLQNYVDRTESRSTEPELIQVKSELPLDP LPVPTEEGNPLLKHYRGPAGDATVASEKESVM

Sequence comparison of the above protein sequences yields the following sequence relationships shown in Table 55B.

10

Table 55B. Comparison of NOV55a against NOV55b.		
Protein Sequence	NOV55a Residues/ Match Residues	Identities/ Similarities for the Matched Region
NOV55b	1..541 28..568	423/541 (78%) 423/541 (78%)

Further analysis of the NOV55a protein yielded the following properties shown in
15 Table 55C.

Table 55C. Protein Sequence Properties NOV55a	
PSort analysis:	0.6000 probability located in plasma membrane; 0.4000 probability located in Golgi body; 0.3000 probability located in endoplasmic reticulum (membrane); 0.3000 probability located in microbody (peroxisome)

SignalP analysis:	Cleavage site between residues 70 and 71

A search of the NOV55a protein against the Geneseq database, a proprietary database that contains sequences published in patents and patent publication, yielded

5   several homologous proteins shown in Table 55D.

Table 55D. Geneseq Results for NOV55a				
Geneseq Identifier	Protein/Organism/Length [Patent #, Date]	NOV55a Residues/ Match Residues	Identities/ Similarities for the Matched Region	Expect Value
ABG61858	Prostate cancer-associated protein #59 - Mammalia, 541 aa. [WO200230268-A2, 18-APR-2002]	1..541 1..541	531/541 (98%) 531/541 (98%)	0.0
AAR95044	Apoptosis participating protein - Homo sapiens, 514 aa. [JP08089257-A, 09-APR-1996]	1..513 1..513	499/513 (97%) 499/513 (97%)	0.0
AAY78144	Human neutral amino acid transporter ASCT1 - Homo sapiens, 532 aa. [US6020479-A, 01-FEB-2000]	32..541 21..532	314/521 (60%) 378/521 (72%)	e-161
AAY99961	Human amino acid transporter ASCT1 protein - Homo sapiens, 532 aa. [US6074828-A, 13-JUN-2000]	32..541 21..532	314/521 (60%) 378/521 (72%)	e-161
AAY97139	ASCT1 human neutral amino acid transporter protein - Homo sapiens, 532 aa. [US6100085-A, 08-AUG-2000]	32..541 21..532	314/521 (60%) 378/521 (72%)	e-161

10   In a BLAST search of public sequence datbases, the NOV55a protein was found to have homology to the proteins shown in the BLASTP data in Table 55E.

Table 55E. Public BLASTP Results for NOV55a

Protein Accession Number	Protein/Organism/Length	NOV55a Residues/ Match Residues	Identities/ Similarities for the Matched Portion	Expect Value
AAD09814	Neutral amino acid transporter - Homo sapiens (Human), 541 aa.	1..541 1..541	532/541 (98%) 532/541 (98%)	0.0
Q15758	Neutral amino acid transporter B(0) (ATB(0)) (Sodium-dependent neutral amino acid transporter type 2) (RD114/simian type D retrovirus receptor) (Baboon M7 virus receptor) - Homo sapiens (Human), 541 aa.	1..541 1..541	531/541 (98%) 531/541 (98%)	0.0
O19105	Neutral amino acid transporter B(0) (ATB(0)) (Sodium-dependent neutral amino acid transporter type 2) - Oryctolagus cuniculus (Rabbit), 541 aa.	1..541 1..541	459/542 (84%) 485/542 (88%)	0.0
Q95JC7	Neutral amino acid transporter B(0) (ATB(0)) (Sodium-dependent neutral amino acid transporter type 2) - Bos taurus (Bovine), 539 aa.	1..541 1..539	465/542 (85%) 486/542 (88%)	0.0
AAM94351	Na+-dependent amino acid transporter ASCT2 - Rattus norvegicus (Rat), 551 aa.	1..541 1..551	445/553 (80%) 471/553 (84%)	0.0

PFam analysis predicts that the NOV55a protein contains the domains shown in the Table 55F.

5

Table 55F. Domain Analysis of NOV55a			
Pfam Domain	NOV55a Match Region	Identities/ Similarities for the Matched Region	Expect Value
SDF	54..485	195/465 (42%) 373/465 (80%)	1.5e-178

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Example B: Sequencing Methodology and Identification of NOVX Clones

1.   **GeneCalling**[TM] **Technology:** This is a proprietary method of performing differential gene expression profiling between two or more samples developed at CuraGen and described by Shimkets, et al., "Gene expression analysis by transcript profiling coupled

5   to a gene database query" Nature Biotechnology 17:198-803 (1999). cDNA was derived from various human samples representing multiple tissue types, normal and diseased states, physiological states, and developmental states from different donors. Samples were obtained as whole tissue, primary cells or tissue cultured primary cells or cell lines. Cells and cell lines may have been treated with biological or chemical agents that regulate gene

10   expression, for example, growth factors, chemokines or steroids. The cDNA thus derived was then digested with up to as many as 120 pairs of restriction enzymes and pairs of linker-adaptors specific for each pair of restriction enzymes were ligated to the appropriate end. The restriction digestion generates a mixture of unique cDNA gene fragments. Limited PCR amplification is performed with primers homologous to the linker adapter

15   sequence where one primer is biotinylated and the other is fluorescently labeled. The doubly labeled material is isolated and the fluorescently labeled single strand is resolved by capillary gel electrophoresis. A computer algorithm compares the electropherograms from an experimental and control group for each of the restriction digestions. This and additional sequence-derived information is used to predict the identity of each differentially expressed

20   gene fragment using a variety of genetic databases. The identity of the gene fragment is confirmed by additional, gene-specific competitive PCR or by isolation and sequencing of the gene fragment.

2.   **SeqCalling**[TM] **Technology:** cDNA was derived from various human samples representing multiple tissue types, normal and diseased states, physiological states,

25   and developmental states from different donors. Samples were obtained as whole tissue, primary cells or tissue cultured primary cells or cell lines. Cells and cell lines may have been treated with biological or chemical agents that regulate gene expression, for example, growth factors, chemokines or steroids. The cDNA thus derived was then sequenced using CuraGen's proprietary SeqCalling technology. Sequence traces were evaluated manually

30   and edited for corrections if appropriate. cDNA sequences from all samples were assembled together, sometimes including public human sequences, using bioinformatic programs to produce a consensus sequence for each assembly. Each assembly is included in CuraGen Corporation's database. Sequences were included as components for assembly

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when the extent of identity with another component was at least 95% over 50 bp. Each assembly represents a gene or portion thereof and includes information on variants, such as splice forms single nucleotide polymorphisms (SNPs), insertions, deletions and other sequence variations.

3. **PathCalling**[TM] **Technology:** The NOVX nucleic acid sequences are derived by laboratory screening of cDNA library by the two-hybrid approach. cDNA fragments covering either the full length of the DNA sequence, or part of the sequence, or both, are sequenced. In silico prediction was based on sequences available in CuraGen Corporation's proprietary sequence databases or in the public human sequence databases, and provided either the full length DNA sequence, or some portion thereof.

The laboratory screening was performed using the methods summarized below:

cDNA libraries were derived from various human samples representing multiple tissue types, normal and diseased states, physiological states, and developmental states from different donors. Samples were obtained as whole tissue, primary cells or tissue cultured primary cells or cell lines. Cells and cell lines may have been treated with biological or chemical agents that regulate gene expression, for example, growth factors, chemokines or steroids. The cDNA thus derived was then directionally cloned into the appropriate two-hybrid vector (Gal4-activation domain (Gal4-AD) fusion). Such cDNA libraries as well as commercially available cDNA libraries from Clontech (Palo Alto, CA) were then transferred from E.coli into a CuraGen Corporation proprietary yeast strain (disclosed in U. S. Patents 6,057,101 and 6,083,693, incorporated herein by reference in their entireties).

Gal4-binding domain (Gal4-BD) fusions of a CuraGen Corportion proprietary library of human sequences was used to screen multiple Gal4-AD fusion cDNA libraries resulting in the selection of yeast hybrid diploids in each of which the Gal4-AD fusion contains an individual cDNA. Each sample was amplified using the polymerase chain reaction (PCR) using non-specific primers at the cDNA insert boundaries. Such PCR product was sequenced; sequence traces were evaluated manually and edited for corrections if appropriate. cDNA sequences from all samples were assembled together, sometimes including public human sequences, using bioinformatic programs to produce a consensus sequence for each assembly. Each assembly is included in CuraGen Corporation's database. Sequences were included as components for assembly when the extent of identity with another component was at least 95% over 50 bp. Each assembly

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represents a gene or portion thereof and includes information on variants, such as splice forms single nucleotide polymorphisms (SNPs), insertions, deletions and other sequence variations.

Physical clone: the cDNA fragment derived by the screening procedure, covering
5　the entire open reading frame is, as a recombinant DNA, cloned into pACT2 plasmid (Clontech) used to make the cDNA library. The recombinant plasmid is inserted into the host and selected by the yeast hybrid diploid generated during the screening procedure by the mating of both CuraGen Corporation proprietary yeast strains N106' and YULH (U. S. Patents 6,057,101 and 6,083,693).

10　　　**4.**　　**RACE:** Techniques based on the polymerase chain reaction such as rapid amplification of cDNA ends (RACE), were used to isolate or complete the predicted sequence of the cDNA of the invention. Usually multiple clones were sequenced from one or more human samples to derive the sequences for fragments. Various human tissue samples from different donors were used for the RACE reaction. The sequences derived
15　from these procedures were included in the SeqCalling Assembly process described in preceding paragraphs.

　　　5.　　**Exon Linking:** The NOVX target sequences identified in the present invention were subjected to the exon linking process to confirm the sequence. PCR primers were designed by starting at the most upstream sequence available, for the forward
20　primer, and at the most downstream sequence available for the reverse primer. In each case, the sequence was examined, walking inward from the respective termini toward the coding sequence, until a suitable sequence that is either unique or highly selective was encountered, or, in the case of the reverse primer, until the stop codon was reached. Such primers were designed based on in silico predictions for the full length cDNA, part (one or
25　more exons) of the DNA or protein sequence of the target sequence, or by translated homology of the predicted exons to closely related human sequences from other species. These primers were then employed in PCR amplification based on the following pool of human cDNAs: adrenal gland, bone marrow, brain - amygdala, brain - cerebellum, brain - hippocampus, brain - substantia nigra, brain - thalamus, brain -whole, fetal brain, fetal
30　kidney, fetal liver, fetal lung, heart, kidney, lymphoma - Raji, mammary gland, pancreas, pituitary gland, placenta, prostate, salivary gland, skeletal muscle, small intestine, spinal cord, spleen, stomach, testis, thyroid, trachea, uterus. Usually the resulting amplicons were gel purified, cloned and sequenced to high redundancy. The PCR product derived from

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exon linking was cloned into the pCR2.1 vector from Invitrogen. The resulting bacterial

clone has an insert covering the entire open reading frame cloned into the pCR2.1 vector.

The resulting sequences from all clones were assembled with themselves, with other

fragments in CuraGen Corporation's database and with public ESTs. Fragments and ESTs

5   were included as components for an assembly when the extent of their identity with another

component of the assembly was at least 95% over 50 bp. In addition, sequence traces were

evaluated manually and edited for corrections if appropriate. These procedures provide the

·sequence reported herein.

6.   **Physical Clone:** Exons were predicted by homology and the intron/exon

10   boundaries were determined using standard genetic rules. Exons were further selected and

refined by means of similarity determination using multiple BLAST (for example, tBlastN,

BlastX, and BlastN) searches, and, in some instances, GeneScan and Grail. Expressed

sequences from both public and proprietary databases were also added when available to

further define and complete the gene sequence. The DNA sequence was then manually

15   corrected for apparent inconsistencies thereby obtaining the sequences encoding the

full-length protein.

The PCR product derived by exon linking, covering the entire open reading frame,

was cloned into the pCR2.1 vector from Invitrogen to provide clones used for expression

and screening purposes.

20   **Example C: Quantitative expression analysis of clones in various cells and tissues**

The quantitative expression of various clones was assessed using microtiter plates

containing RNA samples from a variety of normal and pathology-derived cells, cell lines

and tissues using real time quantitative PCR (RTQ PCR). RTQ PCR was performed on an

Applied Biosystems ABI PRISM® 7700 or an ABI PRISM® 7900 HT Sequence Detection

25   System. Various collections of samples are assembled on the plates, and referred to as

Panel 1 (containing normal tissues and cancer cell lines), Panel 2 (containing samples

derived from tissues from normal and cancer sources), Panel 3 (containing cancer cell

lines), Panel 4 (containing cells and cell lines from normal tissues and cells related to

inflammatory conditions), Panel 5D/5I (containing human tissues and cell lines with an

30   emphasis on metabolic diseases), AI_comprehensive_panel (containing normal tissue and

samples from autoinflammatory diseases), Panel CNSD.01 (containing samples from

normal and diseased brains) and CNS_neurodegeneration_panel (containing samples from

normal and Alzheimer's diseased brains).

RNA integrity from all samples is controlled for quality by visual assessment of
agarose gel electropherograms using 28S and 18S ribosomal RNA staining intensity ratio
as a guide (2:1 to 2.5:1 28s:18s) and the absence of low molecular weight RNAs that would
be indicative of degradation products. Samples are controlled against genomic DNA

5   contamination by RTQ PCR reactions run in the absence of reverse transcriptase using
probe and primer sets designed to amplify across the span of a single exon.

First, the RNA samples were normalized to reference nucleic acids such as
constitutively expressed genes (for example, β-actin and GAPDH). Normalized RNA (5 ul)
was converted to cDNA and analyzed by RTQ-PCR using One Step RT-PCR Master Mix

10   Reagents (Applied Biosystems; Catalog No. 4309169) and gene-specific primers according
to the manufacturer's instructions.

In other cases, non-normalized RNA samples were converted to single strand cDNA
(sscDNA) using Superscript II (Invitrogen Corporation; Catalog No. 18064-147) and
random hexamers according to the manufacturer's instructions. Reactions containing up to

15   10 μg of total RNA were performed in a volume of 20 μl and incubated for 60 minutes at
42°C. This reaction can be scaled up to 50 μg of total RNA in a final volume of 100 μl.
sscDNA samples are then normalized to reference nucleic acids as described previously,
using 1X TaqMan® Universal Master mix (Applied Biosystems; catalog No. 4324020),
following the manufacturer's instructions.

20   Probes and primers were designed for each assay according to Applied Biosystems
Primer Express Software package (version I for Apple Computer's Macintosh Power PC) or
a similar algorithm using the target sequence as input. Default settings were used for
reaction conditions and the following parameters were set before selecting primers: primer
concentration = 250 nM, primer melting temperature (Tm) range = 58°-60°C, primer

25   optimal Tm = 59°C, maximum primer difference = 2°C, probe does not have 5'G, probe Tm
must be 10°C greater than primer Tm, amplicon size 75bp to 100bp. The probes and
primers selected (see below) were synthesized by Synthegen (Houston, TX, USA). Probes
were double purified by HPLC to remove uncoupled dye and evaluated by mass
spectroscopy to verify coupling of reporter and quencher dyes to the 5' and 3' ends of the

30   probe, respectively. Their final concentrations were: forward and reverse primers, 900nM
each, and probe, 200nM.

PCR conditions: When working with RNA samples, normalized RNA from each
tissue and each cell line was spotted in each well of either a 96 well or a 384-well PCR

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plate (Applied Biosystems). PCR cocktails included either a single gene-specific probe and primers set, or two multiplexed probe and primers sets (a set specific for the target clone and another gene-specific set multiplexed with the target probe). PCR reactions were set up using TaqMan® One-Step RT-PCR Master Mix (Applied Biosystems, Catalog No.

5   4313803) following manufacturer's instructions. Reverse transcription was performed at 48°C for 30 minutes followed by amplification/PCR cycles as follows: 95°C 10 min, then 40 cycles of 95°C for 15 seconds, 60°C for 1 minute. Results were recorded as CT values (cycle at which a given sample crosses a threshold level of fluorescence) using a log scale, with the difference in RNA concentration between a given sample and the sample with the

10   lowest CT value being represented as 2 to the power of delta CT. The percent relative expression is then obtained by taking the reciprocal of this RNA difference and multiplying by 100.

When working with sscDNA samples, normalized sscDNA was used as described previously for RNA samples. PCR reactions containing one or two sets of probe and

15   primers were set up as described previously, using 1X TaqMan® Universal Master mix (Applied Biosystems; catalog No. 4324020), following the manufacturer's instructions. PCR amplification was performed as follows: 95°C 10 min, then 40 cycles of 95°C for 15 seconds, 60°C for 1 minute. Results were analyzed and processed as described previously.

Panels 1, 1.1, 1.2, and 1.3D

20   The plates for Panels 1, 1.1, 1.2 and 1.3D include 2 control wells (genomic DNA control and chemistry control) and 94 wells containing cDNA from various samples. The samples in these panels are broken into 2 classes: samples derived from cultured cell lines and samples derived from primary normal tissues. The cell lines are derived from cancers of the following types: lung cancer, breast cancer, melanoma, colon cancer, prostate cancer,

25   CNS cancer, squamous cell carcinoma, ovarian cancer, liver cancer, renal cancer, gastric cancer and pancreatic cancer. Cell lines used in these panels are widely available through the American Type Culture Collection (ATCC), a repository for cultured cell lines, and were cultured using the conditions recommended by the ATCC. The normal tissues found on these panels are comprised of samples derived from all major organ systems from single

30   adult individuals or fetuses. These samples are derived from the following organs: adult skeletal muscle, fetal skeletal muscle, adult heart, fetal heart, adult kidney, fetal kidney, adult liver, fetal liver, adult lung, fetal lung, various regions of the brain, the spleen, bone marrow, lymph node, pancreas, salivary gland, pituitary gland, adrenal gland, spinal cord,

thymus, stomach, small intestine, colon, bladder, trachea, breast, ovary, uterus, placenta,
prostate, testis and adipose.

In the results for Panels 1, 1.1, 1.2 and 1.3D, the following abbreviations are used:

ca. = carcinoma,

* = established from metastasis,

met = metastasis,

s cell var = small cell variant,

non-s = non-sm = non-small,

squam = squamous,

pl. eff = pl effusion = pleural effusion,

glio = glioma,

astro = astrocytoma, and

neuro = neuroblastoma.

General_screening_panel_v1.4, v1.5 and v1.6

The plates for Panels 1.4, 1.5, and 1.6 include 2 control wells (genomic DNA
control and chemistry control) and 94 wells containing cDNA from various samples. The
samples in Panels 1.4, 1.5, and 1.6 are broken into 2 classes: samples derived from cultured
cell lines and samples derived from primary normal tissues. The cell lines are derived from
cancers of the following types: lung cancer, breast cancer, melanoma, colon cancer,
prostate cancer, CNS cancer, squamous cell carcinoma, ovarian cancer, liver cancer, renal
cancer, gastric cancer and pancreatic cancer. Cell lines used in Panels 1.4, 1.5, and 1.6 are
widely available through the American Type Culture Collection (ATCC), a repository for
cultured cell lines, and were cultured using the conditions recommended by the ATCC. The
normal tissues found on Panels 1.4, 1.5, and 1.6 are comprised of pools of samples derived
from all major organ systems from 2 to 5 different adult individuals or fetuses. These
samples are derived from the following organs: adult skeletal muscle, fetal skeletal muscle,
adult heart, fetal heart, adult kidney, fetal kidney, adult liver, fetal liver, adult lung, fetal
lung, various regions of the brain, the spleen, bone marrow, lymph node, pancreas, salivary
gland, pituitary gland, adrenal gland, spinal cord, thymus, stomach, small intestine, colon,
bladder, trachea, breast, ovary, uterus, placenta, prostate, testis and adipose. Abbreviations
are as described for Panels 1, 1.1, 1.2, and 1.3D.

Panels 2D, 2.2, 2.3 and 2.4

The plates for Panels 2D, 2.2, 2.3 and 2.4 generally include 2 control wells and 94 test samples composed of RNA or cDNA isolated from human tissue procured by surgeons working in close cooperation with the National Cancer Institute's Cooperative Human Tissue Network (CHTN) or the National Disease Research Initiative (NDRI) or from

5   Ardais or Clinomics). The tissues are derived from human malignancies and in cases where indicated many malignant tissues have "matched margins" obtained from noncancerous tissue just adjacent to the tumor. These are termed normal adjacent tissues and are denoted "NAT" in the results below. The tumor tissue and the "matched margins" are evaluated by two independent pathologists (the surgical pathologists and again by a pathologist at NDRI/

10   CHTN/Ardais/Clinomics). Unmatched RNA samples from tissues without malignancy (normal tissues) were also obtained from Ardais or Clinomics. This analysis provides a gross histopathological assessment of tumor differentiation grade. Moreover, most samples include the original surgical pathology report that provides information regarding the clinical stage of the patient. These matched margins are taken from the tissue surrounding

15   (i.e. immediately proximal) to the zone of surgery (designated "NAT", for normal adjacent tissue, in Table RR). In addition, RNA and cDNA samples were obtained from various human tissues derived from autopsies performed on elderly people or sudden death victims (accidents, etc.). These tissues were ascertained to be free of disease and were purchased from various commercial sources such as Clontech (Palo Alto, CA), Research Genetics,

20   and Invitrogen.

HASS Panel v 1.0

The HASS panel v 1.0 plates are comprised of 93 cDNA samples and two controls. Specifically, 81 of these samples are derived from cultured human cancer cell lines that had been subjected to serum starvation, acidosis and anoxia for different time periods as well as

25   controls for these treatments, 3 samples of human primary cells, 9 samples of malignant brain cancer (4 medulloblastomas and 5 glioblastomas) and 2 controls. The human cancer cell lines are obtained from ATCC (American Type Culture Collection) and fall into the following tissue groups: breast cancer, prostate cancer, bladder carcinomas, pancreatic cancers and CNS cancer cell lines. These cancer cells are all cultured under standard

30   recommended conditions. The treatments used (serum starvation, acidosis and anoxia) have been previously published in the scientific literature. The primary human cells were obtained from Clonetics (Walkersville, MD) and were grown in the media and conditions recommended by Clonetics. The malignant brain cancer samples are obtained as part of a

collaboration (Henry Ford Cancer Center) and are evaluated by a pathologist prior to

CuraGen receiving the samples . RNA was prepared from these samples using the standard

procedures. The genomic and chemistry control wells have been described previously.

ARDAIS Panel v 1.0

5 The plates for ARDAIS panel v 1.0 generally include 2 control wells and 22 test

samples composed of RNA isolated from human tissue procured by surgeons working in

close cooperation with Ardais Corporation. The tissues are derived from human lung

malignancies (lung adenocarcinoma or lung squamous cell carcinoma) and in cases where

indicated many malignant samples have "matched margins" obtained from noncancerous

10 lung tissue just adjacent to the tumor. These matched margins are taken from the tissue

surrounding (i.e. immediately proximal) to the zone of surgery (designated "NAT", for

normal adjacent tissue) in the results below. The tumor tissue and the "matched margins"

are evaluated by independent pathologists (the surgical pathologists and again by a

pathologist at Ardais). Unmatched malignant and non-malignant RNA samples from lungs

15 were also obtained from Ardais. Additional information from Ardais provides a gross

histopathological assessment of tumor differentiation grade and stage. Moreover, most

samples include the original surgical pathology report that provides information regarding

the clinical state of the patient.

Panel 3D, 3.1 and 3.2

20 The plates of Panel 3D, 3.1, and 3.2 are comprised of 94 cDNA samples and two

control samples. Specifically, 92 of these samples are derived from cultured human cancer

cell lines, 2 samples of human primary cerebellar tissue and 2 controls. The human cell

lines are generally obtained from ATCC (American Type Culture Collection), NCI or the

German tumor cell bank and fall into the following tissue groups: Squamous cell carcinoma

25 of the tongue, breast cancer, prostate cancer, melanoma, epidermoid carcinoma, sarcomas,

bladder carcinomas, pancreatic cancers, kidney cancers, leukemias/lymphomas,

ovarian/uterine/cervical, gastric, colon, lung and CNS cancer cell lines. In addition, there

are two independent samples of cerebellum. These cells are all cultured under standard

recommended conditions and RNA extracted using the standard procedures. The cell lines

30 in panel 3D, 3.1, 3.2, 1, 1.1., 1.2, 1.3D, 1.4, 1.5, and 1.6 are of the most common cell lines

used in the scientific literature.

Panels 4D, 4R, and 4.1D

Panel 4 includes samples on a 96 well plate (2 control wells, 94 test samples)
composed of RNA (Panel 4R) or cDNA (Panels 4D/4.1D) isolated from various human cell
lines or tissues related to inflammatory conditions. Total RNA from control normal tissues
such as colon and lung (Stratagene, La Jolla, CA) and thymus and kidney (Clontech) was

5   employed. Total RNA from liver tissue from cirrhosis patients and kidney from lupus
patients was obtained from BioChain (Biochain Institute, Inc., Hayward, CA). Intestinal
tissue for RNA preparation from patients diagnosed as having Crohn's disease and
ulcerative colitis was obtained from the National Disease Research Interchange (NDRI)
(Philadelphia, PA).

10   Astrocytes, lung fibroblasts, dermal fibroblasts, coronary artery smooth muscle
cells, small airway epithelium, bronchial epithelium, microvascular dermal endothelial
cells, microvascular lung endothelial cells, human pulmonary aortic endothelial cells,
human umbilical vein endothelial cells were all purchased from Clonetics (Walkersville,
MD) and grown in the media supplied for these cell types by Clonetics. These primary cell

15   types were activated with various cytokines or combinations of cytokines for 6 and/or
12-14 hours, as indicated. The following cytokines were used; IL-1 beta at approximately
1-5ng/ml, TNF alpha at approximately 5-10ng/ml, IFN gamma at approximately
20-50ng/ml, IL-4 at approximately 5-10ng/ml, IL-9 at approximately 5-10ng/ml, IL-13 at
approximately 5-10ng/ml. Endothelial cells were sometimes starved for various times by

20   culture in the basal media from Clonetics with 0.1% serum.

Mononuclear cells were prepared from blood of employees at CuraGen
Corporation, using Ficoll. LAK cells were prepared from these cells by culture in DMEM
5% FCS (Hyclone), $100 \mu M$ non essential amino acids (Gibco/Life Technologies,
Rockville, MD), 1mM sodium pyruvate (Gibco), mercaptoethanol $5.5 \times 10^{-5}M$ (Gibco), and

25   10mM Hepes (Gibco) and Interleukin 2 for 4-6 days. Cells were then either activated with
10-20ng/ml PMA and $1-2 \mu g/ml$ ionomycin, IL-12 at 5-10ng/ml, IFN gamma at 20-50ng/ml
and IL-18 at 5-10ng/ml for 6 hours. In some cases, mononuclear cells were cultured for 4-5
days in DMEM 5% FCS (Hyclone), $100 \mu M$ non essential amino acids (Gibco), 1mM
sodium pyruvate (Gibco), mercaptoethanol $5.5 \times 10^{-5}M$ (Gibco), and 10mM Hepes (Gibco)

30   with PHA (phytohemagglutinin) or PWM (pokeweed mitogen) at approximately $5 \mu g/ml$.
Samples were taken at 24, 48 and 72 hours for RNA preparation. MLR (mixed lymphocyte
reaction) samples were obtained by taking blood from two donors, isolating the
mononuclear cells using Ficoll and mixing the isolated mononuclear cells 1:1 at a final

concentration of approximately 2×10^6 cells/ml in DMEM 5% FCS (Hyclone), 100μM non essential amino acids (Gibco), 1mM sodium pyruvate (Gibco), mercaptoethanol (5.5×10^{-5}M) (Gibco), and 10mM Hepes (Gibco). The MLR was cultured and samples taken at various time points ranging from 1- 7 days for RNA preparation.

5   Monocytes were isolated from mononuclear cells using CD14 Miltenyi Beads, +ve VS selection columns and a Vario Magnet according to the manufacturer's instructions. Monocytes were differentiated into dendritic cells by culture in DMEM 5% fetal calf serum (FCS) (Hyclone, Logan, UT), 100μM non essential amino acids (Gibco), 1mM sodium pyruvate (Gibco), mercaptoethanol 5.5×10^{-5}M (Gibco), and 10mM Hepes (Gibco), 50ng/ml

10   GMCSF and 5ng/ml IL-4 for 5-7 days. Macrophages were prepared by culture of monocytes for 5-7 days in DMEM 5% FCS (Hyclone), 100μM non essential amino acids (Gibco), 1mM sodium pyruvate (Gibco), mercaptoethanol 5.5×10^{-5}M (Gibco), 10mM Hepes (Gibco) and 10% AB Human Serum or MCSF at approximately 50ng/ml. Monocytes, macrophages and dendritic cells were stimulated for 6 and 12-14 hours with

15   lipopolysaccharide (LPS) at 100ng/ml. Dendritic cells were also stimulated with anti-CD40 monoclonal antibody (Pharmingen) at 10μg/ml for 6 and 12-14 hours.

CD4 lymphocytes, CD8 lymphocytes and NK cells were also isolated from mononuclear cells using CD4, CD8 and CD56 Miltenyi beads, positive VS selection columns and a Vario Magnet according to the manufacturer's instructions. CD45RA and

20   CD45RO CD4 lymphocytes were isolated by depleting mononuclear cells of CD8, CD56, CD14 and CD19 cells using CD8, CD56, CD14 and CD19 Miltenyi beads and positive selection. CD45RO beads were then used to isolate the CD45RO CD4 lymphocytes with the remaining cells being CD45RA CD4 lymphocytes. CD45RA CD4, CD45RO CD4 and CD8 lymphocytes were placed in DMEM 5% FCS (Hyclone), 100μM non essential amino

25   acids (Gibco), 1mM sodium pyruvate (Gibco), mercaptoethanol 5.5×10^{-5}M (Gibco), and 10mM Hepes (Gibco) and plated at 10^6 cells/ml onto Falcon 6 well tissue culture plates that had been coated overnight with 0.5μg/ml anti-CD28 (Pharmingen) and 3ug/ml anti-CD3 (OKT3, ATCC) in PBS. After 6 and 24 hours, the cells were harvested for RNA preparation. To prepare chronically activated CD8 lymphocytes, we activated the isolated

30   CD8 lymphocytes for 4 days on anti-CD28 and anti-CD3 coated plates and then harvested the cells and expanded them in DMEM 5% FCS (Hyclone), 100μM non essential amino acids (Gibco), 1mM sodium pyruvate (Gibco), mercaptoethanol 5.5×10^{-5}M (Gibco), and 10mM Hepes (Gibco) and IL-2. The expanded CD8 cells were then activated again with

plate bound anti-CD3 and anti-CD28 for 4 days and expanded as before. RNA was isolated 6 and 24 hours after the second activation and after 4 days of the second expansion culture. The isolated NK cells were cultured in DMEM 5% FCS (Hyclone), 100μM non essential amino acids (Gibco), 1mM sodium pyruvate (Gibco), mercaptoethanol 5.5x10⁻⁵M (Gibco),

5   and 10mM Hepes (Gibco) and IL-2 for 4-6 days before RNA was prepared.

To obtain B cells, tonsils were procured from NDRI. The tonsil was cut up with sterile dissecting scissors and then passed through a sieve. Tonsil cells were then spun down and resupended at 10⁶cells/ml in DMEM 5% FCS (Hyclone), 100μM non essential amino acids (Gibco), 1mM sodium pyruvate (Gibco), mercaptoethanol 5.5x10⁻⁵M (Gibco),

10   and 10mM Hepes (Gibco). To activate the cells, we used PWM at 5μg/ml or anti-CD40 (Pharmingen) at approximately 10μg/ml and IL-4 at 5-10ng/ml. Cells were harvested for RNA preparation at 24,48 and 72 hours.

To prepare the primary and secondary Th1/Th2 and Tr1 cells, six-well Falcon plates were coated overnight with 10μg/ml anti-CD28 (Pharmingen) and 2μg/ml OKT3 (ATCC),

15   and then washed twice with PBS. Umbilical cord blood CD4 lymphocytes (Poietic Systems, German Town, MD) were cultured at 10⁵-10⁶cells/ml in DMEM 5% FCS (Hyclone), 100μM non essential amino acids (Gibco), 1mM sodium pyruvate (Gibco), mercaptoethanol 5.5x10⁻⁵M (Gibco), 10mM Hepes (Gibco) and IL-2 (4ng/ml). IL-12 (5ng/ml) and anti-IL4 (1μg/ml) were used to direct to Th1, while IL-4 (5ng/ml) and

20   anti-IFN gamma (1μg/ml) were used to direct to Th2 and IL-10 at 5ng/ml was used to direct to Tr1. After 4-5 days, the activated Th1, Th2 and Tr1 lymphocytes were washed once in DMEM and expanded for 4-7 days in DMEM 5% FCS (Hyclone), 100μM non essential amino acids (Gibco), 1mM sodium pyruvate (Gibco), mercaptoethanol 5.5x10⁻⁵M (Gibco), 10mM Hepes (Gibco) and IL-2 (1ng/ml). Following this, the activated Th1, Th2

25   and Tr1 lymphocytes were re-stimulated for 5 days with anti-CD28/OKT3 and cytokines as described above, but with the addition of anti-CD95L (1μg/ml) to prevent apoptosis. After 4-5 days, the Th1, Th2 and Tr1 lymphocytes were washed and then expanded again with IL-2 for 4-7 days. Activated Th1 and Th2 lymphocytes were maintained in this way for a maximum of three cycles. RNA was prepared from primary and secondary Th1, Th2 and

30   Tr1 after 6 and 24 hours following the second and third activations with plate bound anti-CD3 and anti-CD28 mAbs and 4 days into the second and third expansion cultures in Interleukin 2.

The following leukocyte cells lines were obtained from the ATCC: Ramos, EOL-1, KU-812. EOL cells were further differentiated by culture in 0.1mM dbcAMP at 5×10^5 cells/ml for 8 days, changing the media every 3 days and adjusting the cell concentration to 5×10^5 cells/ml. For the culture of these cells, we used DMEM or RPMI (as
5   recommended by the ATCC), with the addition of 5% FCS (Hyclone), $100 \mu M$ non essential amino acids (Gibco), 1mM sodium pyruvate (Gibco), mercaptoethanol $5.5 \times 10^{-5} M$ (Gibco), 10mM Hepes (Gibco). RNA was either prepared from resting cells or cells activated with PMA at 10ng/ml and ionomycin at $1 \mu g/ml$ for 6 and 14 hours. Keratinocyte line CCD106 and an airway epithelial tumor line NCI-H292 were also obtained from the
10   ATCC. Both were cultured in DMEM 5% FCS (Hyclone), $100 \mu M$ non essential amino acids (Gibco), 1mM sodium pyruvate (Gibco), mercaptoethanol $5.5 \times 10^{-5} M$ (Gibco), and 10mM Hepes (Gibco). CCD1106 cells were activated for 6 and 14 hours with approximately 5 ng/ml TNF alpha and 1ng/ml IL-1 beta, while NCI-H292 cells were activated for 6 and 14 hours with the following cytokines: 5ng/ml IL-4, 5ng/ml IL-9,
15   5ng/ml IL-13 and 25ng/ml IFN gamma.

For these cell lines and blood cells, RNA was prepared by lysing approximately 10^7 cells/ml using Trizol (Gibco BRL). Briefly, 1/10 volume of bromochloropropane (Molecular Research Corporation) was added to the RNA sample, vortexed and after 10 minutes at room temperature, the tubes were spun at 14,000 rpm in a Sorvall SS34 rotor.
20   The aqueous phase was removed and placed in a 15ml Falcon Tube. An equal volume of isopropanol was added and left at -20°C overnight. The precipitated RNA was spun down at 9,000 rpm for 15 min in a Sorvall SS34 rotor and washed in 70% ethanol. The pellet was redissolved in $300 \mu l$ of RNAse-free water and $35 \mu l$ buffer (Promega) $5 \mu l$ DTT, $7 \mu l$ RNAsin and $8 \mu l$ DNAse were added. The tube was incubated at 37°C for 30 minutes to
25   remove contaminating genomic DNA, extracted once with phenol chloroform and re-precipitated with 1/10 volume of 3M sodium acetate and 2 volumes of 100% ethanol. The RNA was spun down and placed in RNAse free water. RNA was stored at -80°C.

AI_comprehensive panel_v1.0

The plates for AI_comprehensive panel_v1.0 include two control wells and 89 test
30   samples comprised of cDNA isolated from surgical and postmortem human tissues obtained from the Backus Hospital and Clinomics (Frederick, MD). Total RNA was extracted from tissue samples from the Backus Hospital in the Facility at CuraGen. Total RNA from other tissues was obtained from Clinomics.

Joint tissues including synovial fluid, synovium, bone and cartilage were obtained
from patients undergoing total knee or hip replacement surgery at the Backus Hospital.
Tissue samples were immediately snap frozen in liquid nitrogen to ensure that isolated
RNA was of optimal quality and not degraded. Additional samples of osteoarthritis and
5   rheumatoid arthritis joint tissues were obtained from Clinomics. Normal control tissues
were supplied by Clinomics and were obtained during autopsy of trauma victims.

Surgical specimens of psoriatic tissues and adjacent matched tissues were provided
as total RNA by Clinomics. Two male and two female patients were selected between the
ages of 25 and 47. None of the patients were taking prescription drugs at the time samples
10   were isolated.

Surgical specimens of diseased colon from patients with ulcerative colitis and
Crohns disease and adjacent matched tissues were obtained from Clinomics. Bowel tissue
from three female and three male Crohn's patients between the ages of 41-69 were used.
Two patients were not on prescription medication while the others were taking
15   dexamethasone, phenobarbital, or tylenol. Ulcerative colitis tissue was from three male and
four female patients. Four of the patients were taking lebvid and two were on
phenobarbital.

Total RNA from post mortem lung tissue from trauma victims with no disease or
with emphysema, asthma or COPD was purchased from Clinomics. Emphysema patients
20   ranged in age from 40-70 and all were smokers, this age range was chosen to focus on
patients with cigarette-linked emphysema and to avoid those patients with
alpha-1anti-trypsin deficiencies. Asthma patients ranged in age from 36-75, and excluded
smokers to prevent those patients that could also have COPD. COPD patients ranged in age
from 35-80 and included both smokers and non-smokers. Most patients were taking
25   corticosteroids, and bronchodilators.

In the labels employed to identify tissues in the AI_comprehensive panel_v1.0
panel, the following abbreviations are used:

AI = Autoimmunity

Syn = Synovial

30   Normal = No apparent disease

Rep22 /Rep20 = individual patients

RA = Rheumatoid arthritis

Backus = From Backus Hospital

333

OA = Osteoarthritis

(SS) (BA) (MF) = Individual patients

Adj = Adjacent tissue

Match control = adjacent tissues

5   -M = Male

-F = Female

COPD = Chronic obstructive pulmonary disease

AI.05 chondrosarcoma

10

The AI.05 chondrosarcoma plates are comprised of SW1353 cells that had been subjected
to serum starvation, and treatment with cytokines that are known to induce MMP (1, 3 and 13)
synthesis (eg. IL1beta). These treatments include: IL-1β (10 ng/ml), IL-1β + TNF-α (50 ng/ml),
IL-1β + Oncostatin (50 ng/ml) and PMA (100 ng/ml). The SW1353 cells were obtained from
15   ATCC (American Type Culture Collection) and were all cultured under standard recommended
conditions. The SW1353 cells were plated at 3 x10^5 cells/ml (in DMEM medium-10 % FBS)
in 6-well plate. The treatment was done in triplicate, for 6 and 18 h. The supernatants were
collected for analysis of MMP 1, 3 and 13 production and for RNA extraction. RNA was prepared
from these samples using the standard procedures.

20   ### Panels 5D and 5I

The plates for Panel 5D and 5I include two control wells and a variety of cDNAs
isolated from human tissues and cell lines with an emphasis on metabolic diseases.
Metabolic tissues were obtained from patients enrolled in the Gestational Diabetes study.
Cells were obtained during different stages in the differentiation of adipocytes from human
25   mesenchymal stem cells. Human pancreatic islets were also obtained.

In the Gestational Diabetes study subjects are young (18 - 40 years), otherwise
healthy women with and without gestational diabetes undergoing routine (elective)
Caesarean section. After delivery of the infant, when the surgical incisions were being
repaired/closed, the obstetrician removed a small sample (<1 cc) of the exposed metabolic
30   tissues during the closure of each surgical level. The biopsy material was rinsed in sterile
saline, blotted and fast frozen within 5 minutes from the time of removal. The tissue was
then flash frozen in liquid nitrogen and stored, individually, in sterile screw-top tubes and
kept on dry ice for shipment to or to be picked up by CuraGen. The metabolic tissues of

interest include uterine wall (smooth muscle), visceral adipose, skeletal muscle (rectus) and subcutaneous adipose. Patient descriptions are as follows:

Patient 2: Diabetic Hispanic, overweight, not on insulin

Patient 7-9: Nondiabetic Caucasian and obese (BMI>30)

5   Patient 10: Diabetic Hispanic, overweight, on insulin

Patient 11: Nondiabetic African American and overweight

Patient 12: Diabetic Hispanic on insulin

Adiocyte differentiation was induced in donor progenitor cells obtained from Osirus (a division of Clonetics/BioWhittaker) in triplicate, except for Donor 3U which had only 10   two replicates. Scientists at Clonetics isolated, grew and differentiated human mesenchymal stem cells (HuMSCs) for CuraGen based on the published protocol found in Mark F. Pittenger, et al., Multilineage Potential of Adult Human Mesenchymal Stem Cells Science Apr 2 1999: 143-147. Clonetics provided Trizol lysates or frozen pellets suitable for mRNA isolation and ds cDNA production. A general description of each donor is as 15   follows:

Donor 2 and 3 U: Mesenchymal Stem cells, Undifferentiated Adipose

Donor 2 and 3 AM: Adipose, AdiposeMidway Differentiated

Donor 2 and 3 AD: Adipose, Adipose Differentiated

Human cell lines were generally obtained from ATCC (American Type Culture 20   Collection), NCI or the German tumor cell bank and fall into the following tissue groups: kidney proximal convoluted tubule, uterine smooth muscle cells, small intestine, liver HepG2 cancer cells, heart primary stromal cells, and adrenal cortical adenoma cells. These cells are all cultured under standard recommended conditions and RNA extracted using the standard procedures. All samples were processed at CuraGen to produce single stranded 25   cDNA.

Panel 5I contains all samples previously described with the addition of pancreatic islets from a 58 year old female patient obtained from the Diabetes Research Institute at the University of Miami School of Medicine. Islet tissue was processed to total RNA at an outside source and delivered to CuraGen for addition to panel 5I.

30   In the labels employed to identify tissues in the 5D and 5I panels, the following abbreviations are used:

335

GO Adipose = Greater Omentum Adipose

SK = Skeletal Muscle

UT = Uterus

PL = Placenta

5   AD = Adipose Differentiated

AM = Adipose Midway Differentiated

U = Undifferentiated Stem Cells

Panel CNSD.01

The plates for Panel CNSD.01 include two control wells and 94 test samples

10   comprised of cDNA isolated from postmortem human brain tissue obtained from the

Harvard Brain Tissue Resource Center. Brains are removed from calvaria of donors

between 4 and 24 hours after death, sectioned by neuroanatomists, and frozen at -80°C in

liquid nitrogen vapor. All brains are sectioned and examined by neuropathologists to

confirm diagnoses with clear associated neuropathology.

15   Disease diagnoses are taken from patient records. The panel contains two brains

from each of the following diagnoses: Alzheimer's disease, Parkinson's disease,

Huntington's disease, Progressive Supernuclear Palsy, Depression, and "Normal controls".

Within each of these brains, the following regions are represented: cingulate gyrus,

temporal pole, globus palladus, substantia nigra, Brodman Area 4 (primary motor strip),

20   Brodman Area 7 (parietal cortex), Brodman Area 9 (prefrontal cortex), and Brodman area

17 (occipital cortex). Not all brain regions are represented in all cases; e.g., Huntington's

disease is characterized in part by neurodegeneration in the globus palladus, thus this

region is impossible to obtain from confirmed Huntington's cases. Likewise Parkinson's

disease is characterized by degeneration of the substantia nigra making this region more

25   difficult to obtain. Normal control brains were examined for neuropathology and found to

be free of any pathology consistent with neurodegeneration.

In the labels employed to identify tissues in the CNS panel, the following

abbreviations are used:

PSP = Progressive supranuclear palsy

30   Sub Nigra = Substantia nigra

Glob Palladus= Globus palladus

Temp Pole = Temporal pole

Cing Gyr = Cingulate gyrus

BA 4 = Brodman Area 4

Panel CNS_Neurodegeneration_V1.0

The plates for Panel CNS_Neurodegeneration_V1.0 include two control wells and

5   47 test samples comprised of cDNA isolated from postmortem human brain tissue obtained

from the Harvard Brain Tissue Resource Center (McLean Hospital) and the Human Brain

and Spinal Fluid Resource Center (VA Greater Los Angeles Healthcare System). Brains are

removed from calvaria of donors between 4 and 24 hours after death, sectioned by

neuroanatomists, and frozen at -80°C in liquid nitrogen vapor. All brains are sectioned and

10   examined by neuropathologists to confirm diagnoses with clear associated neuropathology.

Disease diagnoses are taken from patient records. The panel contains six brains

from Alzheimer's disease (AD) patients, and eight brains from "Normal controls" who

showed no evidence of dementia prior to death. The eight normal control brains are divided

into two categories: Controls with no dementia and no Alzheimer's like pathology

15   (Controls) and controls with no dementia but evidence of severe Alzheimer's like

pathology, (specifically senile plaque load rated as level 3 on a scale of 0-3; 0 = no

evidence of plaques, 3 = severe AD senile plaque load). Within each of these brains, the

following regions are represented: hippocampus, temporal cortex (Brodman Area 21),

parietal cortex (Brodman area 7), and occipital cortex (Brodman area 17). These regions

20   were chosen to encompass all levels of neurodegeneration in AD. The hippocampus is a

region of early and severe neuronal loss in AD; the temporal cortex is known to show

neurodegeneration in AD after the hippocampus; the parietal cortex shows moderate

neuronal death in the late stages of the disease; the occipital cortex is spared in AD and

therefore acts as a "control" region within AD patients. Not all brain regions are

25   represented in all cases.

In the labels employed to identify tissues in the CNS_Neurodegeneration_V1.0

panel, the following abbreviations are used:

AD = Alzheimer's disease brain; patient was demented and showed AD-like

pathology upon autopsy

30   Control = Control brains; patient not demented, showing no neuropathology

Control (Path) = Control brains; pateint not demented but showing sever AD-like

pathology

337

SupTemporal Ctx = Superior Temporal Cortex

Inf Temporal Ctx = Inferior Temporal Cortex

A. CG106764-01: RHO/RAC-INTERACTING CITRON KINASE.

5   Expression of gene CG106764-01 was assessed using the primer-probe set Ag2100, described in Table AA. Results of the RTQ-PCR runs are shown in Tables AB, AC, AD, AE, AF, AG, AH and AI.

Table AA. Probe Name Ag2100

10

Primers	Sequence	Length	Start Position	SEQ ID No
Forward	5'-agatccctggaacagaggatt-3'	21	2446	249
Probe	TET-5'-tgtctgaagccaataaacttgcagca-3'-TAMRA	26	2474	250
Reverse	5'-ccttcatgttcctttgggtaa-3'	21	2513	251

Table AB. AI.05 chondrosarcoma

15

Tissue Name	Rel. Exp.(%) g2100, Run 306913849	Tissue Name	Rel. Exp.(%) Ag2100, Run 306913849
138353_PMA (18hrs)	9.3	138346_IL-1beta + Oncostatin M (6hrs)	64.2
138352_IL-1beta + Oncostatin M (18hrs)	5.5	138345_IL-1beta+TNFa (6hrs)	44.8
138351_IL-1beta+TNFa (18hrs)	12.5	138344_IL-1beta (6hrs)	25.5
138350_IL-1beta (18hrs)	12.5	138349_Untreated-serum starved (6hrs)	100.0
138354_Untreated-complete medium (18hrs)	13.2	138348_Untreated-complete medium (6hrs)	41.2
138347_PMA (6hrs)	34.9		

Table AC. AI comprehensive panel v1.0

Tissue Name	Rel. Exp.(%) Ag2100, Run 211059880	Rel. Exp.(%) Ag2100, Run 212328504	issue Name	Rel. Exp.(%) Ag2100, Run 211059880	Rel. Exp.(%) Ag2100, Run 212328504
110967 COPD-F	0.5	0.8	112427 Match Control Psoriasis-F	2.9	1.8
110980 COPD-F	1.5	1.2	112418 Psoriasis-M	0.8	0.8
110968 COPD-M	0.4	0.6	112723 Match Control Psoriasis-M	6.1	7.4
110977 COPD-M	1.5	1.9	112419 Psoriasis-M	1.0	1.3
110989 Emphysema-F	4.2	6.0	112424 Match Control Psoriasis-M	0.4	1.2
110992 Emphysema-F	2.8	2.9	112420 Psoriasis-M	1.8	2.4
110993 Emphysema-F	0.9	0.8	112425 Match Control Psoriasis-M	2.2	2.7
110994 Emphysema-F	0.7	0.4	104689 (MF) OA Bone-Backus	12.1	13.2
110995 Emphysema-F	2.0	5.4	104690 (MF) Adj "Normal" Bone-Backus	5.4	4.2
110996 Emphysema-F	2.2	2.4	104691 (MF) OA Synovium-Backus	43.2	35.6
110997 Asthma-M	1.9	3.1	104692 (BA) OA Cartilage-Backus	0.9	0.4
111001 Asthma-F	1.4	2.7	104694 (BA) OA Bone-Backus	16.8	16.7
111002 Asthma-F	1.0	1.0	104695 (BA) Adj "Normal" Bone-Backus	6.5	6.1
111003 Atopic Asthma-F	4.0	2.2	104696 (BA) OA Synovium-Backus	24.0	24.1
111004 Atopic Asthma-F	16.6	17.0	104700 (SS) OA Bone-Backus	12.2	35.1
111005 Atopic Asthma-F	7.2	5.5	104701 (SS) Adj "Normal" Bone-Backus	7.9	9.5
111006 Atopic Asthma-F	0.9	0.7	104702 (SS) OA Synovium-Backus	8.2	7.9
111417 Allergy-M	1.9	2.4	117093 OA Cartilage Rep7	2.0	2.3
112347 Allergy-M	0.0	0.1	112672 OA Bone5	1.9	0.8
112349 Normal Lung-F	0.0	0.0	112673 OA Synovium5	0.3	1.2

112357 Normal Lung-F	6.1	6.0	112674 OA Synovial Fluid cells5	0.5	0.4
112354 Normal Lung-M	1.5	2.3	117100 OA Cartilage Rep14	0.4	0.3
112374 Crohns-F	2.9	5.2	112756 OA Bone9	**100.0**	**100.0**
112389 Match Control Crohns-F	9.0	6.8	112757 OA Synovium9	0.5	0.2
112375 Crohns-F	2.5	3.8	112758 OA Synovial Fluid Cells9	0.8	1.5
112732 Match Control Crohns-F	3.8	5.4	117125 RA Cartilage Rep2	1.0	0.6
112725 Crohns-M	0.1	0.7	113492 Bone2 RA	2.8	3.6
112387 Match Control Crohns-M	1.0	1.4	113493 Synovium2 RA	1.7	0.7
112378 Crohns-M	0.0	0.0	113494 Syn Fluid Cells RA	0.9	2.1
112390 Match Control Crohns-M	2.5	1.8	113499 Cartilage4 RA	2.1	1.8
112726 Crohns-M	3.8	5.9	113500 Bone4 RA	1.8	2.5
112731 Match Control Crohns-M	3.6	6.7	113501 Synovium4 RA	2.1	2.3
112380 Ulcer Col-F	4.9	4.9	113502 Syn Fluid Cells4 RA	1.0	0.8
112734 Match Control Ulcer Col-F	12.6	12.0	113495 Cartilage3 RA	2.5	2.6
112384 Ulcer Col-F	6.6	10.2	113496 Bone3 RA	2.0	2.1
112737 Match Control Ulcer Col-F	4.2	6.1	113497 Synovium3 RA	1.4	1.4
112386 Ulcer Col-F	0.5	1.2	113498 Syn Fluid Cells3 RA	2.9	3.2
112738 Match Control Ulcer Col-F	7.5	7.9	117106 Normal Cartilage Rep20	0.1	0.7
112381 Ulcer Col-M	0.1	0.1	113663 Bone3 Normal	0.3	0.1
112735 Match Control Ulcer Col-M	2.9	2.3	113664 Synovium3 Normal	0.0	0.0
112382 Ulcer Col-M	6.7	8.4	113665 Syn Fluid Cells3 Normal	0.1	0.2
112394 Match Control Ulcer Col-M	0.5	0.5	117107 Normal Cartilage Rep22	0.9	0.3
112383 Ulcer Col-M	12.1	14.6	113667 Bone4 Normal	0.4	0.7

| 112736 Match Control Ulcer Col-M | 3.5 | 5.3 | 113668 Synovium4 Normal | 1.0 | 1.1 |
| 112423 Psoriasis-F | 1.4 | 1.1 | 113669 Syn Fluid Cells4 Normal | 1.0 | 0.7 |

Table AD. CNS_neurodegeneration_v1.0

Tissue Name	Rel. Exp.(%) Ag2100, Run 207929343	Tissue Name	Rel. Exp.(%) Ag2100, Run 207929343
AD 1 Hippo	5.2	Control (Path) 3 Temporal Ctx	8.5
AD 2 Hippo	9.3	Control (Path) 4 Temporal Ctx	55.5
AD 3 Hippo	6.7	AD 1 Occipital Ctx	31.6
AD 4 Hippo	7.2	AD 2 Occipital Ctx (Missing)	0.0
AD 5 Hippo	100.0	AD 3 Occipital Ctx	8.4
AD 6 Hippo	16.5	AD 4 Occipital Ctx	28.7
Control 2 Hippo	17.7	AD 5 Occipital Ctx	52.5
Control 4 Hippo	3.4	AD 6 Occipital Ctx	22.8
Control (Path) 3 Hippo	4.4	Control 1 Occipital Ctx	3.9
AD 1 Temporal Ctx	15.7	Control 2 Occipital Ctx	64.6
AD 2 Temporal Ctx	26.4	Control 3 Occipital Ctx	40.6
AD 3 Temporal Ctx	12.3	Control 4 Occipital Ctx	6.4
AD 4 Temporal Ctx	24.3	Control (Path) 1 Occipital Ctx	77.9
AD 5 Inf Temporal Ctx	65.5	Control (Path) 2 Occipital Ctx	28.5
AD 5 Sup Temporal Ctx	20.9	Control (Path) 3 Occipital Ctx	1.5
AD 6 Inf Temporal Ctx	44.1	Control (Path) 4 Occipital Ctx	40.9
AD 6 Sup Temporal Ctx	59.0	Control 1 Parietal Ctx	7.8
Control 1 Temporal Ctx	9.5	Control 2 Parietal Ctx	34.4
Control 2 Temporal Ctx	34.6	Control 3 Parietal Ctx	15.8
Control 3 Temporal Ctx	0.0	Control (Path) 1 Parietal Ctx	68.8
Control 3 Temporal Ctx	10.4	Control (Path) 2 Parietal Ctx	32.3
Control (Path) 1 Temporal Ctx	68.8	Control (Path) 3 Parietal Ctx	4.9
Control (Path) 2 Temporal Ctx	49.7	Control (Path) 4 Parietal Ctx	58.6

Table AE. Panel 1.3D

Tissue Name	Rel. Exp. %) Ag2100, Run 152517508	Tissue Name	Rel. Exp.(%) Ag2100, Run 152517508
Liver adenocarcinoma	11.7	Kidney (fetal)	1.8
Pancreas	0.0	Renal ca. 786-0	7.1
Pancreatic ca. CAPAN 2	3.2	Renal ca. A498	3.7
Adrenal gland	1.4	Renal ca. RXF 393	3.1
Thyroid	0.1	Renal ca. ACHN	4.4
Salivary gland	0.1	Renal ca. UO-31	6.3
Pituitary gland	2.1	Renal ca. TK-10	3.2
Brain (fetal)	2.1	Liver	0.0
Brain (whole)	24.7	Liver (fetal)	3.8
Brain (amygdala)	11.2	Liver ca. (hepatoblast) HepG2	3.2
Brain (cerebellum)	2.7	Lung	0.3
Brain (hippocampus)	36.3	Lung (fetal)	0.9
Brain (substantia nigra)	1.5	Lung ca. (small cell) LX-1	6.6
Brain (thalamus)	30.4	Lung ca. (small cell) NCI-H69	8.5
Cerebral Cortex	100.0	Lung ca. (s.cell var.) SHP-77	7.5
Spinal cord	2.5	Lung ca. (large cell)NCI-H460	0.0
glio/astro U87-MG	6.4	Lung ca. (non-sm. cell) A549	0.2
glio/astro U-118-MG	33.7	Lung ca. (non-s.cell) NCI-H23	10.4
astrocytoma SW1783	5.9	Lung ca. (non-s.cell) HOP-62	1.4
neuro*; met SK-N-AS	14.5	Lung ca. (non-s.cl) NCI-H522	5.3
astrocytoma SF-539	7.4	Lung ca. (squam.) SW 900	3.2
astrocytoma SNB-75	5.8	Lung ca. (squam.) NCI-H596	7.2
glioma SNB-19	1.0	Mammary gland	0.2
glioma·U251	2.4	Breast ca.* (pl.ef) MCF-7	5.6
glioma SF-295	0.9	Breast ca.* (pl.ef) MDA-MB-231	14.5
Heart (fetal)	0.4	Breast ca.* (pl.ef) T47D	2.4
Heart	0.1	Breast ca. BT-549	6.8
Skeletal muscle (fetal)	3.4	Breast ca. MDA-N	14.0
Skeletal muscle	0.1	Ovary	2.2
Bone marrow	5.4	Ovarian ca. OVCAR-3	2.5
Thymus	2.1	Ovarian ca. OVCAR-4	0.8
Spleen	0.6	Ovarian ca. OVCAR-5	2.7
Lymph node	0.4	Ovarian ca. OVCAR-8	3.2
Colorectal	1.8	Ovarian ca. IGROV-1	2.0
Stomach	1.0	Ovarian ca.* (ascites) SK-OV-3	7.4
Small intestine	1.6	Uterus	0.0
Colon ca. SW480	13.1	Placenta	0.2

Colon ca.* SW620(SW480 met)	4.5	Prostate	0.2
Colon ca. HT29	4.1	Prostate ca.* (bone met)PC-3	2.0
Colon ca. HCT-116	5.0	Testis	4.0
Colon ca. CaCo-2	5.9	Melanoma Hs688(A).T	0.7
Colon ca. tissue(ODO3866)	2.8	Melanoma* (met) Hs688(B).T	0.3
Colon ca. HCC-2998	3.7	Melanoma UACC-62	0.5
Gastric ca.* (liver met) NCI-N87	2.3	Melanoma M14	7.2
Bladder	0.9	Melanoma LOX IMVI	2.8
Trachea	0.7	Melanoma* (met) SK-MEL-5	5.8
Kidney	0.7	Adipose	0.2

Table AF. Panel 2.2

5

Tissue Name	Rel. Ep.(%) Ag2100, Run 174166901	Tissue Name	Rel. Exp.(%) Ag2100, Run 174166901
Normal Colon	6.3	Kidney Margin (OD04348)	30.4
Colon cancer (OD06064)	13.4	Kidney malignant cancer (OD06204B)	3.6
Colon Margin (OD06064)	9.0	Kidney normal adjacent tissue (OD06204E)	10.5
Colon cancer (OD06159)	4.5	Kidney Cancer (OD04450-01)	2.4
Colon Margin (OD06159)	5.9	Kidney Margin (OD04450-03)	13.3
Colon cancer (OD06297-04)	3.8	Kidney Cancer 8120613	6.7
Colon Margin (OD06297-05)	9.9	Kidney Margin 8120614	1.2
CC Gr.2 ascend colon (ODO3921)	4.4	Kidney Cancer 9010320	1.7
CC Margin (ODO3921)	2.8	Kidney Margin 9010321	4.5
Colon cancer metastasis (OD06104)	1.7	Kidney Cancer 8120607	0.5
Lung Margin (OD06104)	3.1	Kidney Margin 8120608	1.7
Colon mets to lung (OD04451-01)	9.6	Normal Uterus	1.1
Lung Margin (OD04451-02)	3.2	Uterine Cancer 064011	1.5
Normal Prostate	1.2	Normal Thyroid	0.0
Prostate Cancer (OD04410)	0.0	Thyroid Cancer 064010	0.6
Prostate Margin (OD04410)	0.7	Thyroid Cancer A302152	5.3
Normal Ovary	2.8	Thyroid Margin A302153	0.0
Ovarian cancer (OD06283-03)	11.7	Normal Breast	3.0
Ovarian Margin (OD06283-07)	3.0	Breast Cancer (OD04566)	8.1
Ovarian Cancer 064008	1.1	Breast Cancer 1024	2.9
Ovarian cancer (OD06145)	0.9	Breast Cancer (OD04590-01)	14.8
Ovarian Margin (OD06145)	0.0	Breast Cancer Mets (OD04590-03)	3.2

Ovarian cancer (OD06455-03)	15.8	Breast Cancer Metastasis (OD04655-05)	5.4
Ovarian Margin (OD06455-07)	1.8	Breast Cancer 064006	3.1
Normal Lung	1.2	Breast Cancer 9100266	2.6
Invasive poor diff. lung adeno (OD04945-01	8.4	Breast Margin 9100265	2.3
Lung Margin (OD04945-03)	1.2	Breast Cancer A209073	1.8
Lung Malignant Cancer (OD03126)	5.0	Breast Margin A2090734	2.5
Lung Margin (OD03126)	0.6	Breast cancer (OD06083)	17.1
Lung Cancer (OD05014A)	10.2	Breast cancer node metastasis (OD06083)	14.7
Lung Margin (OD05014B)	9.0	Normal Liver	0.4
Lung cancer (OD06081)	10.1	Liver Cancer 1026	0.0
Lung Margin (OD06081)	4.0	Liver Cancer 1025	1.8
Lung Cancer (OD04237-01)	4.1	Liver Cancer 6004-T	1.1
Lung Margin (OD04237-02)	2.0	Liver Tissue 6004-N	2.5
Ocular Melanoma Metastasis	0.9	Liver Cancer 6005-T	1.6
Ocular Melanoma Margin (Liver)	0.4	Liver Tissue 6005-N	0.0
Melanoma Metastasis	10.4	Liver Cancer 064003	0.7
Melanoma Margin (Lung)	2.0	Normal Bladder	2.9
Normal Kidney	5.0	Bladder Cancer 1023	1.5
Kidney Ca, Nuclear grade 2 (OD04338)	15.4	Bladder Cancer A302173	17.8
Kidney Margin (OD04338)	5.0	Normal Stomach	10.4
Kidney Ca Nuclear grade 1/2 (OD04339)	100.0	Gastric Cancer 9060397	1.1
Kidney Margin (OD04339)	9.3	Stomach Margin 9060396	0.7
Kidney Ca, Clear cell type (OD04340)	14.0	Gastric Cancer 9060395	2.8
Kidney Margin (OD04340)	11.3	Stomach Margin 9060394	2.8
Kidney Ca, Nuclear grade 3 (OD04348)	9.0	Gastric Cancer 064005	6.0

Table AG. Panel 3D

5

Tissue Name	Rel. Exp (%) Ag2100, Run 164796104	Tissue Name	Rel. Exp.(%) Ag2100, Run 164796104
Daoy- Medulloblastoma	7.3	Ca Ski- Cervical epidermoid carcinoma (metastasis)	21.0
TE671- Medulloblastoma	3.8	ES-2- Ovarian clear cell carcinoma	11.7

D283 Med- Medulloblastoma	15.7	Ramos- Stimulated with PMA/ionomycin 6h	10.8
PFSK-1- Primitive Neuroectodermal	11.2	Ramos- Stimulated with PMA/ionomycin 14h	6.2
XF-498- CNS	21.2	MEG-01- Chronic myelogenous leukemia (megokaryoblast)	5.8
SNB-78- Glioma	11.3	Raji- Burkitt's lymphoma	6.7
SF-268- Glioblastoma	7.6	Daudi- Burkitt's lymphoma	14.8
T98G- Glioblastoma	12.0	U266- B-cell plasmacytoma	5.1
SK-N-SH- Neuroblastoma (metastasis)	5.6	CA46- Burkitt's lymphoma	5.0
SF-295- Glioblastoma	12.4	RL- non-Hodgkin's B-cell lymphoma	3.8
Cerebellum	16.2	JM1- pre-B-cell lymphoma	11.5
Cerebellum	3.6	Jurkat- T cell leukemia	12.5
NCI-H292- Mucoepidermoid lung carcinoma	14.0	TF-1- Erythroleukemia	9.9
DMS-114- Small cell lung cancer	10.4	HUT 78- T-cell lymphoma	14.7
DMS-79- Small cell lung cancer	100.0	U937- Histiocytic lymphoma	8.1
NCI-H146- Small cell lung cancer	14.3	KU-812- Myelogenous leukemia	17.7
NCI-H526- Small cell lung cancer	19.8	769-P- Clear cell renal carcinoma	6.3
NCI-N417- Small cell lung cancer	5.8	Caki-2- Clear cell renal carcinoma	9.5
NCI-H82- Small cell lung cancer	10.2	SW 839- Clear cell renal carcinoma	5.2
NCI-H157- Squamous cell lung cancer (metastasis)	13.8	G401- Wilms' tumor	6.3
NCI-H1155- Large cell lung cancer	36.1	Hs766T- Pancreatic carcinoma (LN metastasis)	15.7
NCI-H1299- Large cell lung cancer	22.7	CAPAN-1- Pancreatic adenocarcinoma (liver metastasis)	8.6
NCI-H727- Lung carcinoid	14.4	SU86.86- Pancreatic carcinoma (liver metastasis)	14.1
NCI-UMC-11- Lung carcinoid	25.9	BxPC-3- Pancreatic adenocarcinoma	9.4
LX-1- Small cell lung cancer	11.0	HPAC- Pancreatic adenocarcinoma	14.5
Colo-205- Colon cancer	12.7	MIA PaCa-2- Pancreatic carcinoma	2.6
KM12- Colon cancer	17.2	CFPAC-1- Pancreatic ductal adenocarcinoma	38.7
KM20L2- Colon cancer	7.0	PANC-1- Pancreatic epithelioid ductal carcinoma	19.5
NCI-H716- Colon cancer	19.5	T24- Bladder carcinma (transitional cell)	9.0
SW-48- Colon adenocarcinoma	10.6	5637- Bladder carcinoma	10.5
SW1116- Colon adenocarcinoma	7.7	HT-1197- Bladder carcinoma	4.8

LS 174T- Colon adenocarcinoma	9.8	UM-UC-3- Bladder carcinoma (transitional cell)	13.3
SW-948- Colon adenocarcinoma	1.4	A204- Rhabdomyosarcoma	15.2
SW-480- Colon adenocarcinoma	7.6	HT-1080- Fibrosarcoma	11.9
NCI-SNU-5- Gastric carcinoma	14.9	MG-63- Osteosarcoma	7.3
KATO III- Gastric carcinoma	18.8	SK-LMS-1- Leiomyosarcoma (vulva)	48.0
NCI-SNU-16- Gastric carcinoma	12.6	SJRH30- Rhabdomyosarcoma (met to bone marrow)	10.2
NCI-SNU-1- Gastric carcinoma	12.3	A431- Epidermoid carcinoma	12.2
RF-1- Gastric adenocarcinoma	5.3	WM266-4- Melanoma	21.9
RF-48- Gastric adenocarcinoma	7.6	DU 145- Prostate carcinoma (brain metastasis)	0.2
MKN-45- Gastric carcinoma	11.7	MDA-MB-468- Breast adenocarcinoma	5.6
NCI-N87- Gastric carcinoma	9.3	SCC-4- Squamous cell carcinoma of tongue	0.3
OVCAR-5- Ovarian carcinoma	3.0	SCC-9- Squamous cell carcinoma of tongue	0.3
RL95-2- Uterine carcinoma	4.5	SCC-15- Squamous cell carcinoma of tongue	0.2
HelaS3- Cervical adenocarcinoma	9.0	CAL 27- Squamous cell carcinoma of tongue	19.9

Table AH. Panel 4D

5

Tissue Name	Rel. Exp (%) Ag2100, Run 152800279	Tissue Name	Rel. Exp.(%) Ag2100, Run 152800279
Secondary Th1 act	15.4	HUVEC IL-1beta	12.2
Secondary Th2 act	11.9	HUVEC IFN gamma	16.6
Secondary Tr1 act	15.6	HUVEC TNF alpha + IFN gamma	11.8
Secondary Th1 rest	4.9	HUVEC TNF alpha + IL4	11.4
Secondary Th2 rest	3.3	HUVEC IL-11	8.2
Secondary Tr1 rest	6.0	Lung Microvascular EC none	7.3
Primary Th1 act	13.6	Lung Microvascular EC TNFalpha + IL-1beta	6.3
Primary Th2 act	12.0	Microvascular Dermal EC none	23.3
Primary Tr1 act	22.2	Microsvasular Dermal EC TNFalpha + IL-1beta	10.5
Primary Th1 rest	**100.0**	Bronchial epithelium TNFalpha + IL1beta	0.6
Primary Th2 rest	37.9	Small airway epithelium none	1.6

Primary Tr1 rest	29.3	Small airway epithelium TNFalpha + IL-1beta	7.4
CD45RA CD4 lymphocyte act	13.6	Coronery artery SMC rest	4.4
CD45RO CD4 lymphocyte act	15.4	Coronery artery SMC TNFalpha + IL-1beta	2.0
CD8 lymphocyte act	10.6	Astrocytes rest	1.3
Secondary CD8 lymphocyte rest	7.9	Astrocytes TNFalpha + IL-1beta	0.5
Secondary CD8 lymphocyte act	17.3	KU-812 (Basophil) rest	22.4
CD4 lymphocyte none	0.5	KU-812 (Basophil) PMA/ionomycin	28.5
2ry Th1/Th2/Tr1_anti-CD95 CH11	17.1	CCD1106 (Keratinocytes) none	14.3
LAK cells rest	3.6	CCD1106 (Keratinocytes) TNFalpha + IL-1beta	18.4
LAK cells IL-2	16.8	Liver cirrhosis	0.5
LAK cells IL-2+IL-12	8.4	Lupus kidney	3.3
LAK cells IL-2+IFN gamma	16.4	NCI-H292 none	29.5
LAK cells IL-2+ IL-18	16.8	NCI-H292 IL-4	27.7
LAK cells PMA/ionomycin	0.6	NCI-H292 IL-9	32.3
NK Cells IL-2 rest	15.3	NCI-H292 IL-13	13.4
Two Way MLR 3 day	1.8	NCI-H292 IFN gamma	11.0
Two Way MLR 5 day	6.1	HPAEC none	8.5
Two Way MLR 7 day	10.1	HPAEC TNF alpha + IL-1 beta	7.7
PBMC rest	0.1	Lung fibroblast none	6.3
PBMC PWM	25.5	Lung fibroblast TNF alpha + IL-1 beta	9.0
PBMC PHA-L	24.0	Lung fibroblast IL-4	3.7
Ramos (B cell) none	17.7	Lung fibroblast IL-9	5.0
Ramos (B cell) ionomycin	92.0	Lung fibroblast IL-13	1.7
B lymphocytes PWM	48.6	Lung fibroblast IFN gamma	3.4
B lymphocytes CD40L and IL-4	16.4	Dermal fibroblast CCD1070 rest	57.4
EOL-1 dbcAMP	10.5	Dermal fibroblast CCD1070 TNF alpha	79.0
EOL-1 dbcAMP PMA/ionomycin	7.0	Dermal fibroblast CCD1070 IL-1 beta	21.8
Dendritic cells none	0.5	Dermal fibroblast IFN gamma	22.2
Dendritic cells LPS	0.0	Dermal fibroblast IL-4	45.7
Dendritic cells anti-CD40	0.0	IBD Colitis 2	0.9
Monocytes rest	0.2	IBD Crohn's	1.0
Monocytes LPS	0.0	Colon	3.7
Macrophages rest	4.4	Lung	1.5
Macrophages LPS	0.6	Thymus	13.0
HUVEC none	24.7	Kidney	31.2
HUVEC starved	43.5		

Table AI. Panel CNS_1

Tissue Name	Rel Exp.(%) Ag2100, Run 171649357	Tissue Name	Rel. Exp.(%) Ag2100, Run 171649357
BA4 Control	23.8	BA17 PSP	35.4
BA4 Control2	19.1	BA17 PSP2	18.3
BA4 Alzheimer's2	7.3	Sub Nigra Control	11.6
BA4 Parkinson's	43.8	Sub Nigra Control2	5.0
BA4 Parkinson's2	60.7	Sub Nigra Alzheimer's2	4.6
BA4 Huntington's	23.3	Sub Nigra Parkinson's2	11.8
BA4 Huntington's2	14.7	Sub Nigra Huntington's	16.0
BA4 PSP	13.8	Sub Nigra Huntington's2	8.8
BA4 PSP2	26.2	Sub Nigra PSP2	1.7
BA4 Depression	15.4	Sub Nigra Depression	2.7
BA4 Depression2	17.0	Sub Nigra Depression2	8.0
BA7 Control	36.6	Glob Palladus Control	8.4
BA7 Control2	17.4	Glob Palladus Control2	10.8
BA7 Alzheimer's2	11.3	Glob Palladus Alzheimer's	1.8
BA7 Parkinson's	21.9	Glob Palladus Alzheimer's2	8.3
BA7 Parkinson's2	36.1	Glob Palladus Parkinson's	51.1
BA7 Huntington's	56.3	Glob Palladus Parkinson's2	12.9
BA7 Huntington's2	45.1	Glob Palladus PSP	9.3
BA7 PSP	44.4	Glob Palladus PSP2	9.9
BA7 PSP2	17.6	Glob Palladus Depression	6.0
BA7 Depression	8.5	Temp Pole Control	9.8
BA9 Control	31.9	Temp Pole Control2	21.5
BA9 Control2	34.4	Temp Pole Alzheimer's	6.6
BA9 Alzheimer's	8.0	Temp Pole Alzheimer's2	8.1
BA9 Alzheimer's2	20.0	Temp Pole Parkinson's	33.0
BA9 Parkinson's	40.6	Temp Pole Parkinson's2	24.8
BA9 Parkinson's2	31.4	Temp Pole Huntington's	33.2
BA9 Huntington's	41.5	Temp Pole PSP	8.8
BA9 Huntington's2	21.8	Temp Pole PSP2	6.0
BA9 PSP	17.8	Temp Pole Depression2	17.0
BA9 PSP2	8.2	Cing Gyr Control	23.3
BA9 Depression	10.5	Cing Gyr Control2	17.8
BA9 Depression2	16.2	Cing Gyr Alzheimer's	7.3
BA17 Control	58.2	Cing Gyr Alzheimer's2	10.4
BA17 Control2	41.8	Cing Gyr Parkinson's	13.4
BA17 Alzheimer's2	27.0	Cing Gyr Parkinson's2	17.0
BA17 Parkinson's	58.6	Cing Gyr Huntington's	28.3

BA17 Parkinson's2	69.3	Cing Gyr Huntington's2	10.6
BA17 Huntington's	44.4	Cing Gyr PSP	7.2
BA17 Huntington's2	31.9	Cing Gyr PSP2	4.0
BA17 Depression	13.6	Cing Gyr Depression	6.9
BA17 Depression2	100.0	Cing Gyr Depression2	10.4

AI.05 chondrosarcoma Summary: Ag2100 Highest expression of this gene is detected in untreated serum starved chondrosarcoma cell line (SW1353) (CT=27). Interestingly, expression of this gene appears to be somewhat down regulated upon IL-1 treatment, a potent activator of pro-inflammatory cytokines and matrix metalloproteinases which participate in the destruction of cartilage observed in Osteoarthritis (OA). Modulation of the expression of this transcript in chondrocytes by either small molecules or antisense might be important for preventing the degeneration of cartilage observed in OA

AI_comprehensive panel_v1.0 Summary: Ag2100 Highest expression of this gene is detected in osteoarthritis (OA) bone (CTs=27-28). This gene is highly expressed in bone isolated from 5 different osteoarthritic (OA) patients, synovium in 3 out of 5 OA patients, but not in cartilege from OA patients nor in any tissues from rheumatoid arthritis (RA) patients or control samples. Thus, small molecule therapeutics designed against the protein encoded for by this gene could reduce or inhibit inflammation. Anti-sense therapeutics that would block the translation of the transcript and protein production could also inhibit inflammatory processes. These types of therapeutics could be important in the treatment of diseases such as osteoarthritis

CNS_neurodegeneration_v1.0 Summary: Ag2100 This panel confirms the expression of this gene at low levels in the brains of an independent group of individuals. However, no differential expression of this gene was detected between Alzheimer's diseased postmortem brains and those of non-demented controls in this experiment. Please see Panel 1.3D for a discussion of this gene in treatment of central nervous system disorders.

Panel 1.3D Summary: Ag2100 Expression of this gene is highest in cerebral cortex (CT = 26.3). This gene is expressed at moderate levels in all the regions of the CNS including amygdala, cerebellum, hippocampus, substantia nigra, thalamus, spinal cord, and fetal brain. This gene encodes a protein with homology to citron-kinase. Citron-kinase (Citron-K) has been proposed by in vitro studies to be a crucial effector of Rho in regulation of cytokinesis. Citron-K is essential for cytokinesis in vivo in specific neuronal

precursors and may play a fundamental role in specific human malformative syndromes of the CNS (Di Cunto et al., 2000, Neuron 28:115-127, PMID: 11086988). General inhibitors of the RHO/RAC-INTERACTING CITRON KINASE family disrupt endothelial tight junctions, suggesting that specific modulators of this brain-preferential family member could be useful in delivery of therapeutics across the blood brain barrier. These general inhibitors also influence intracellular calcium flux, which is a central component of many important neuronal processes, such as apoptosis, neurotransmitter release and signal transduction (Jezior et al., 2001, Br. J. Pharmacol. 134:78-87, PMID: 11522599; Walsh et al., 2001, Gastroenterology 121:566-579, PMID: 11522741). Thus, modulators of the function of the protein encoded by this gene may prove useful in the treatment of neurodegenerative disorders involving apoptosis, such as spinal muscular atrophy, Alzheimer's disease, Huntington's disease, Parkinson's disease, and others. Diseases involving neurotransmitters or signal transduction, such as schizophrenia, mania, stroke, epilepsy and depression may also benefit from agents that modulate the function of the this gene product.

This gene also shows moderate to low expression in several metabolic tissues including adrenal gland, pituitary gland, gastrointestinal tract, fetal heart, fetal skeletal muscle and fetal liver. Therefore, therapeutic modulation of the activity of this gene may prove useful in the treatment of endocrine/metabolically related diseases, such as obesity and diabetes.

Interestingly, expression of this gene is higher in fetal tissues (CTs=31) as compared to the corresponding adult liver, and skeletal muscle (CTs=37-40). This observation suggests that expression of this gene can be used to distinguish fetal from adult liver and skeletal muscle. In addition, the relative overexpression of this gene in fetal tissue suggests that the protein product may enhance liver and muscle growth or development in the fetus and thus may also act in a regenerative capacity in the adult. Therefore, therapeutic modulation of the protein encoded by this gene could be useful in treatment of liver and skeletal muscle related diseases.

Moderate levels of expression of this gene is also seen in cluster of cancer cell lines derived from pancreatic, gastric, colon, lung, liver, renal, breast, ovarian, prostate, melanoma and brain cancers. Thus, therapeutic modulation of the expression or function of this gene may be effective in the treatment of pancreatic, gastric, colon, lung, liver, renal, breast, ovarian, prostate, melanoma and brain cancers.

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Panel 2.2 Summary: Ag2100 Expression of this gene is highest in a kidney cancer sample (CT=28). In addition, significant expression of this gene is also seen in a number of normal and cancer tissues including colon, lung, ovary, breast, kidney, thyroid, liver, bladder, and stomach. Interestingly, this gene is expressed at slightly higher levels in most of the tumors than in the normal matched tissue. Thus, expression of this gene could be used to distinguish between cancerous tissue and normal tissue. In addition, therapeutic modulation of this gene product, through the use of small molecule drugs or antibodies, might be of benefit in the treatment of cancer.

Panel 3D Summary: Ag2100 Expression of this gene is highest in a lung cancer cell line (CT = 26). However, low to moderate expression is also seen in the majority of cancer cell lines on this panel, suggesting that this gene may play an important role in many cell types.

Panel 4D Summary: Ag2100 Highest expression of this gene is detected in resting primary Th1 cells (CT=24.5). Moderate to low levels of expression of this gene is seen in members of the T-cell, B-cell, endothelial cell, macrophage/monocyte, and peripheral blood mononuclear cell family, as well as epithelial and fibroblast cell types from lung and skin, and normal tissues represented by colon, lung, thymus and kidney. Interestingly, this gene is highly induced in Ramos B cells treated with PMA and ionomycin, in non-transformed B cells and PBMC treated with PWM. All three of these observations are consistent with this gene being induced in B cells after activation. This gene product has homology to the RHO/RAC-interacting citron kinase. Thus citron kinase encoded by this gene may play an important role in T cell activation, by regulating TCR-mediated T cell spreading, chemotaxis and other chemokine responses and in apoptosis. Likewise, this putative kinase may also be important in B cell motility, antigen receptor mediated activation and apoptosis.

Small molecule therapeutics designed against the protein encoded for by this gene could reduce or inhibit inflammation. Anti-sense therapeutics that would block the translation of the transcript and protein production could also inhibit inflammatory processes. These types of therapeutics could be important in the treatment of diseases such as osteoarthritis. Likewise, these therapeutics could be important in the treatment of asthma, psoriasis, diabetes, and IBD, which require activated T cells, as well as diseases that involve B cell activation such as systemic lupus erythematosus.

Panel CNS_1 Summary: Ag2100 This panel confirms the expression of this gene at low levels in the brains of an independent group of individuals. Please see Panel 1.3D for a discussion of this gene in treatment of central nervous system disorders.

B. CG117662-02: Renal renin precursor like.

Expression of gene CG117662-02 was assessed using the primer-probe sets Ag2078 and Ag5185, described in Tables BA and BB. Results of the RTQ-PCR runs are shown in Tables BC, BD, BE, BF and BG.

Table BA. Probe Name Ag2078

Primers	Sequence	Length	Start Position	SEQ ID No
Forward	5'-accttcaaagtcgtctttgaca-3'	22	292	252
Probe	TET-5'-ctccaagtgcagccgtctctacactg-3'-TAMRA	26	342	253
Reverse	5'-cgaagagcttgtgatacacaca-3'	22	370	254

Table BB. Probe Name Ag5185

Primers	Sequence	Length	Start Position	SEQ ID No
Forward	5'-ccgtgtctgtggggtcat-3'	18	491	255
Probe	TET-5'-attggtagacaccggtgcatcctaca-3'-TAMRA	26	540	256
Reverse	5'-tggagctggtagaacctgaga-3'	21	566	257

Table BC. CNS_neurodegeneration_v1.0

Tissue Name	Rel. Exp.(%) Ag5185, Run 226559655	issue Name	Rel. Exp.(%) Ag5185, Run 226559655
AD 1 Hippo	5.7	Control (Path) 3 Temporal Ctx	48.6
AD 2 Hippo	82.4	Control (Path) 4 Temporal Ctx	54.3
AD 3 Hippo	11.4	AD 1 Occipital Ctx	12.2
AD 4 Hippo	50.0	AD 2 Occipital Ctx (Missing)	0.0
AD 5 Hippo	22.5	AD 3 Occipital Ctx	18.8

AD 6 Hippo	15.2	AD 4 Occipital Ctx	19.8
Control 2 Hippo	9.6	AD 5 Occipital Ctx	12.1
Control 4 Hippo	18.3	AD 6 Occipital Ctx	25.0
Control (Path) 3 Hippo	85.3	Control 1 Occipital Ctx	26.2
AD 1 Temporal Ctx	38.4	Control 2 Occipital Ctx	3.6
AD 2 Temporal Ctx	74.7	Control 3 Occipital Ctx	40.6
AD 3 Temporal Ctx	0.0	Control 4 Occipital Ctx	20.9
AD 4 Temporal Ctx	49.0	Control (Path) 1 Occipital Ctx	39.2
AD 5 Inf Temporal Ctx	31.6	Control (Path) 2 Occipital Ctx	18.3
AD 5 Sup Temporal Ctx	36.3	Control (Path) 3 Occipital Ctx	0.0
AD 6 Inf Temporal Ctx	55.5	Control (Path) 4 Occipital Ctx	0.0
AD 6 Sup Temporal Ctx	63.3	Control 1 Parietal Ctx	46.7
Control 1 Temporal Ctx	**100.0**	Control 2 Parietal Ctx	0.0
Control 2 Temporal Ctx	40.6	Control 3 Parietal Ctx	12.2
Control 3 Temporal Ctx	47.0	Control (Path) 1 Parietal Ctx	65.5
Control 3 Temporal Ctx	24.7	Control (Path) 2 Parietal Ctx	23.8
Control (Path) 1 Temporal Ctx	50.7	Control (Path) 3 Parietal Ctx	0.0
Control (Path) 2 Temporal Ctx	65.5	Control (Path) 4 Parietal Ctx	57.4

Table BD. General screening panel v1.5

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Tissue Name	Rel. Exp.(%) Ag5185, Run 228757766	issue Name	Rel. Exp.(%) Ag5185, Run 228757766
Adipose	1.0	Renal ca. TK-10	0.0
Melanoma* Hs688(A).T	0.2	Bladder	0.5
Melanoma* Hs688(B).T	0.1	Gastric ca. (liver met.) NCI-N87	1.1
Melanoma* M14	0.1	Gastric ca. KATO III	0.3
Melanoma* LOXIMVI	0.1	Colon ca. SW-948	18.2
Melanoma* SK-MEL-5	0.2	Colon ca. SW480	0.6
Squamous cell carcinoma SCC-4	0.4	Colon ca.* (SW480 met) SW620	0.5
Testis Pool	8.4	Colon ca. HT29	1.6
Prostate ca.* (bone met) PC-3	1.5	Colon ca. HCT-116	0.5
Prostate Pool	0.6	Colon ca. CaCo-2	0.2
Placenta	3.0	Colon cancer tissue	2.6
Uterus Pool	1.5	Colon ca. SW1116	0.1
Ovarian ca. OVCAR-3	0.9	Colon ca. Colo-205	0.0
Ovarian ca. SK-OV-3	0.2	Colon ca. SW-48	0.8
Ovarian ca. OVCAR-4	0.7	Colon Pool	4.7
Ovarian ca. OVCAR-5	4.7	Small Intestine Pool	4.0
Ovarian ca. IGROV-1	0.0	Stomach Pool	2.3

Ovarian ca. OVCAR-8	0.2	Bone Marrow Pool	2.5
Ovary	6.7	Fetal Heart	0.2
Breast ca. MCF-7	0.5	Heart Pool	1.6
Breast ca. MDA-MB-231	0.6	Lymph Node Pool	12.6
Breast ca. BT 549	0.2	Fetal Skeletal Muscle	0.3
Breast ca. T47D	2.1	Skeletal Muscle Pool	0.4
Breast ca. MDA-N	0.0	Spleen Pool	0.1
Breast Pool	5.0	Thymus Pool	3.8
Trachea	1.0	CNS cancer (glio/astro) U87-MG	0.0
Lung	22.1	CNS cancer (glio/astro) U-118-MG	0.1
Fetal Lung	0.6	CNS cancer (neuro;met) SK-N-AS	0.0
Lung ca. NCI-N417	0.4	CNS cancer (astro) SF-539	0.0
Lung ca. LX-1	0.3	CNS cancer (astro) SNB-75	0.0
Lung ca. NCI-H146	0.0	CNS cancer (glio) SNB-19	0.3
Lung ca. SHP-77	0.1	CNS cancer (glio) SF-295	0.5
Lung ca. A549	0.0	Brain (Amygdala) Pool	0.4
Lung ca. NCI-H526	0.5	Brain (cerebellum)	0.5
Lung ca. NCI-H23	1.4	Brain (fetal)	0.0
Lung ca. NCI-H460	2.0	Brain (Hippocampus) Pool	0.2
Lung ca. HOP-62	0.1	Cerebral Cortex Pool	0.3
Lung ca. NCI-H522	0.6	Brain (Substantia nigra) Pool	0.3
Liver	1.0	Brain (Thalamus) Pool	0.6
Fetal Liver	1.0	Brain (whole)	0.8
Liver ca. HepG2	0.0	Spinal Cord Pool	0.5
Kidney Pool	4.2	Adrenal Gland	2.6
Fetal Kidney	100.0	Pituitary gland Pool	0.6
Renal ca. 786-0	0.0	Salivary Gland	0.5
Renal ca. A498	0.0	Thyroid (female)	0.1
Renal ca. ACHN	0.2	Pancreatic ca. CAPAN2	0.2
Renal ca. UO-31	0.3	Pancreas Pool	4.9

Table BE. Panel 1.3D

5

Tissue Name	Rel. Exp.(%) g2078, Run 165626684	Rel. Exp.(%) Ag2078, Run 165627496	Rel. Exp.(%) Ag2078, Run 165678122	Tissue Name	Rel. Exp.(%) Ag2078, Run 165626684	Rel. Exp.(%) Ag2078, Run 165627496	Rel. Exp.(%) Ag2078, Run 165678122
Liver adenocarcinoma	0.0	0.1	0.1	Kidney (fetal)	100.0	100.0	100.0
Pancreas	0.0	0.0	0.0	Renal ca. 786-0	0.0	0.0	0.0
Pancreatic ca. CAPAN 2	0.0	0.0	0.2	Renal ca. A498	0.0	0.0	0.1

Adrenal gland	0.5	0.5	0.3	Renal ca. RXF 393	0.0	0.0	0.0
Thyroid	0.0	0.0	0.0	Renal ca. ACHN	0.0	0.0	0.0
Salivary gland	0.0	0.1	0.0	Renal ca. UO-31	0.0	0.0	0.1
Pituitary gland	0.0	0.2	0.0	Renal ca. TK-10	0.0	0.1	0.0
Brain (fetal)	0.0	0.0	0.0	Liver	0.3	0.3	0.0
Brain (whole)	0.0	0.0	0.1	Liver (fetal)	0.6	0.7	0.5
Brain (amygdala)	0.1	0.0	0.0	Liver ca. (hepatoblast) HepG2	0.0	0.0	0.0
Brain (cerebellum)	0.1	0.0	0.1	Lung	0.0	0.0	0.1
Brain (hippocampus)	0.0	0.3	0.0	Lung (fetal)	0.1	0.1	0.0
Brain (substantia nigra)	0.0	0.0	0.1	Lung ca. (small cell) LX-1	0.0	0.0	0.0
Brain (thalamus)	0.1	0.0	0.1	Lung ca. (small cell) NCI-H69	0.0	0.0	0.0
Cerebral Cortex	0.0	0.0	0.2	Lung ca. (s.cell var.) SHP-77	0.0	0.1	0.0
Spinal cord	0.0	0.0	0.0	Lung ca. (large cell)NCI-H460	0.0	0.0	0.0
glio/astro U87-MG	0.0	0.0	0.0	Lung ca. (non-sm. cell) A549	0.0	0.0	0.0
glio/astro U-118-MG	0.0	0.0	0.0	Lung ca. (non-s.cell) NCI-H23	0.0	0.0	0.0
astrocytoma SW1783	0.0	0.0	0.1	Lung ca. (non-s.cell) HOP-62	0.1	0.0	0.0
neuro*; met SK-N-AS	0.0	0.1	0.0	Lung ca. (non-s.cl) NCI-H522	0.0	0.0	0.0
astrocytoma SF-539	0.0	0.0	0.0	Lung ca. (squam.) SW 900	0.1	0.1	0.0
astrocytoma SNB-75	0.0	0.0	0.2	Lung ca. (squam.) NCI-H596	0.0	0.0	0.0
glioma SNB-19	0.0	0.0	0.0	Mammary gland	0.2	0.2	0.1
glioma U251	0.0	0.0	0.0	Breast ca.* (pl.ef) MCF-7	0.0	0.0	0.1

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glioma SF-295	0.0	0.0	0.0	Breast ca.* (pl.ef) MDA-MB-231	0.1	0.0	0.0
Heart (fetal)	0.0	0.0	0.0	Breast ca.* (pl.ef) T47D	0.1	0.0	0.0
Heart	0.0	0.0	0.0	Breast ca. BT-549	0.0	0.0	0.0
Skeletal muscle (fetal)	0.0	0.0	0.0	Breast ca. MDA-N	0.0	0.0	0.0
Skeletal muscle	0.0	0.0	0.0	Ovary	0.6	0.8	0.6
Bone marrow	0.0	0.0	0.0	Ovarian ca. OVCAR-3	0.1	0.1	0.0
Thymus	0.0	0.0	0.0	Ovarian ca. OVCAR-4	0.0	0.1	0.0
Spleen	0.0	0.0	0.0	Ovarian ca. OVCAR-5	0.2	0.2	0.1
Lymph node	0.0	0.1	0.0	Ovarian ca. OVCAR-8	0.0	0.0	0.0
Colorectal	0.0	0.0	0.0	Ovarian ca. IGROV-1	0.0	0.0	0.0
Stomach	0.0	0.0	0.1	Ovarian ca.* (ascites) SK-OV-3	0.0	0.0	0.0
Small intestine	0.1	0.0	0.0	Uterus	1.7	1.1	1.1
Colon ca. SW480	0.0	0.0	0.0	Placenta	0.7	1.2	0.7
Colon ca.* SW620(SW480 met)	0.0	0.0	0.0	Prostate	0.1	0.0	0.1
Colon ca. HT29	0.2	0.3	0.3	Prostate ca.* (bone met)PC-3	0.2	0.2	0.0
Colon ca. HCT-116	0.0	0.0	0.0	Testis	0.2	0.1	0.2
Colon ca. CaCo-2	0.0	0.0	0.0	Melanoma Hs688(A).T	0.0	0.0	0.0
Colon ca. tissue(ODO386 6)	0.2	0.1	0.5	Melanoma* (met) Hs688(B).T	0.0	0.0	0.0
Colon ca. HCC-2998	0.1	0.3	0.1	Melanoma UACC-62	0.0	0.0	0.0
Gastric ca.* (liver met) NCI-N87	0.0	0.1	0.0	Melanoma M14	0.0	0.0	0.0
Bladder	0.0	0.0	0.0	Melanoma LOX IMVI	0.0	0.0	0.1
Trachea	0.1	0.0	0.0	Melanoma* (met) SK-MEL-5	0.0	0.0	0.0
Kidney	11.2	10.8	8.7	Adipose	0.0	0.2	0.0

Table BF. Panel 4D

5

Tissue Name	Rel. xp.(%) Ag2078, Run 161905846	Tissue Name	Rel. Exp.(%) Ag2078, Run 161905846
Secondary Th1 act	0.0	HUVEC IL-1beta	0.0
Secondary Th2 act	0.0	HUVEC IFN gamma	0.0
Secondary Tr1 act	0.0	HUVEC TNF alpha + IFN gamma	0.0
Secondary Th1 rest	0.0	HUVEC TNF alpha + IL4	0.0
Secondary Th2 rest	0.0	HUVEC IL-11	0.0
Secondary Tr1 rest	0.0	Lung Microvascular EC none	0.2
Primary Th1 act	0.0	Lung Microvascular EC TNFalpha + IL-1beta	0.2
Primary Th2 act	0.0	Microvascular Dermal EC none	0.3
Primary Tr1 act	0.0	Microsvasular Dermal EC TNFalpha + IL-1beta	0.1
Primary Th1 rest	0.0	Bronchial epithelium TNFalpha + IL1beta	0.1
Primary Th2 rest	0.0	Small airway epithelium none	0.0
Primary Tr1 rest	0.0	Small airway epithelium TNFalpha + IL-1beta	0.5
CD45RA CD4 lymphocyte act	0.8	Coronery artery SMC rest	0.1
CD45RO CD4 lymphocyte act	0.0	Coronery artery SMC TNFalpha + IL-1beta	0.1
CD8 lymphocyte act	0.0	Astrocytes rest	0.0
Secondary CD8 lymphocyte rest	0.0	Astrocytes TNFalpha + IL-1beta	0.1
Secondary CD8 lymphocyte act	0.0	KU-812 (Basophil) rest	0.0
CD4 lymphocyte none	0.0	KU-812 (Basophil) PMA/ionomycin	0.0
2ry Th1/Th2/Tr1_anti-CD95 CH11	0.0	CCD1106 (Keratinocytes) none	0.0
LAK cells rest	0.0	CCD1106 (Keratinocytes) TNFalpha + IL-1beta	0.0
LAK cells IL-2	0.0	Liver cirrhosis	0.4
LAK cells IL-2+IL-12	0.0	Lupus kidney	3.9
LAK cells IL-2+IFN gamma	0.0	NCI-H292 none	1.3
LAK cells IL-2+ IL-18	0.0	NCI-H292 IL-4	0.5
LAK cells PMA/ionomycin	0.1	NCI-H292 IL-9	1.9
NK Cells IL-2 rest	0.0	NCI-H292 IL-13	0.3
Two Way MLR 3 day	0.0	NCI-H292 IFN gamma	1.0
Two Way MLR 5 day	0.0	HPAEC none	0.0
Two Way MLR 7 day	0.0	HPAEC TNF alpha + IL-1 beta	0.0

PBMC rest	0.1	Lung fibroblast none	0.0
PBMC PWM	0.0	Lung fibroblast TNF alpha + IL-1 beta	0.0
PBMC PHA-L	0.0	Lung fibroblast IL-4	0.0
Ramos (B cell) none	0.0	Lung fibroblast IL-9	0.0
Ramos (B cell) ionomycin	0.0	Lung fibroblast IL-13	0.0
B lymphocytes PWM	0.0	Lung fibroblast IFN gamma	0.0
B lymphocytes CD40L and IL-4	0.0	Dermal fibroblast CCD1070 rest	5.9
EOL-1 dbcAMP	0.0	Dermal fibroblast CCD1070 TNF alpha	4.5
EOL-1 dbcAMP PMA/ionomycin	0.2	Dermal fibroblast CCD1070 IL-1 beta	3.1
Dendritic cells none	0.0	Dermal fibroblast IFN gamma	0.0
Dendritic cells LPS	0.0	Dermal fibroblast IL-4	0.0
Dendritic cells anti-CD40	0.0	IBD Colitis 2	0.0
Monocytes rest	0.0	IBD Crohn's	0.0
Monocytes LPS	0.0	Colon	0.0
Macrophages rest	0.0	Lung	0.2
Macrophages LPS	0.0	Thymus	100.0
HUVEC none	0.4	Kidney	0.4
HUVEC starved	0.2		

Table BG. Panel 5D

5

Tissue Name	Rel. Ex.(%) Ag2078, Run 168095527	Tissue Name	Rel. Exp.(%) Ag2078, Run 168095527
97457_Patient-02go_adipose	11.7	94709_Donor 2 AM - A_adipose	0.0
97476_Patient-07sk_skeletal muscle	0.0	94710_Donor 2 AM - B_adipose	0.0
97477_Patient-07ut_uterus	2.8	94711_Donor 2 AM - C_adipose	0.0
97478_Patient-07pl_placenta	12.9	94712_Donor 2 AD - A_adipose	1.0
97481_Patient-08sk_skeletal muscle	0.0	94713_Donor 2 AD - B_adipose	0.0
97482_Patient-08ut_uterus	22.8	94714_Donor 2 AD - C_adipose	0.0
97483_Patient-08pl_placenta	4.5	94742_Donor 3 U - A_Mesenchymal Stem Cells	0.0
97486_Patient-09sk_skeletal muscle	0.0	94743_Donor 3 U - B_Mesenchymal Stem Cells	0.0
97487_Patient-09ut_uterus	0.0	94730_Donor 3 AM - A_adipose	0.9
97488_Patient-09pl_placenta	2.7	94731_Donor 3 AM - B_adipose	0.0
97492_Patient-10ut_uterus	100.0	94732_Donor 3 AM - C_adipose	0.0

97493_Patient-10pl_placenta	5.4	94733_Donor 3 AD - A_adipose	0.0
97495_Patient-11go_adipose	6.0	94734_Donor 3 AD - B_adipose	0.0
97496_Patient-11sk_skeletal muscle	0.0	94735_Donor 3 AD - C_adipose	0.0
97497_Patient-11ut_uterus	12.8	77138_Liver_HepG2untreated	0.0
97498_Patient-11pl_placenta	8.5	73556_Heart_Cardiac stromal cells (primary)	0.0
97500_Patient-12go_adipose	87.1	81735_Small Intestine	0.0
97501_Patient-12sk_skeletal muscle	0.0	72409_Kidney_Proximal Convoluted Tubule	0.0
97502_Patient-12ut_uterus	4.6	82685_Small intestine_Duodenum	0.0
97503_Patient-12pl_placenta	8.0	90650_Adrenal_Adrenocortical adenoma	0.0
94721_Donor 2 U - A_Mesenchymal Stem Cells	0.0	72410_Kidney_HRCE	1.1
94722_Donor 2 U - B_Mesenchymal Stem Cells	0.0	72411_Kidney_HRE	5.3
94723_Donor 2 U - C_Mesenchymal Stem Cells	0.0	73139_Uterus_Uterine smooth muscle cells	2.4

CNS_neurodegeneration_v1.0 Summary: Ag5185 Low levels of expression of this gene is seen in control temporal cortex and in a hippocampus sample from an Alzheimer patient (CTs=34.6-34.9). Therefore, therapeutic modulation of this gene may be useful in the neurological disorders including seizure and memory related diseases.

General_screening_panel_v1.5 Summary: Ag5185 Highest expression of this gene is detected in fetal kidney (CT=26.7). Interestingly, expression of this gene is higher in fetal as compared to adult kidney (CT=31). This observation suggests that expression of this gene can be used to distinguish fetal from adult kidney and also from other samples in this panel. In addition, the relative overexpression of this gene in fetal tissue suggests that the protein product may enhance kidney growth or development in the fetus and thus may also act in a regenerative capacity in the adult. Therefore, therapeutic modulation of the protein encoded by this gene could be useful in treatment of kidney related diseases including lupus and glomerulonephritis.

Moderate to low levels of expression of this gene is also seen in tissues with metabolic/endocrine functions such as pancreas, adiposes, adrenal and pituitary glands, heart, skeletal muscle, and gastrointestinal tract. Therefore, therapeutic modulation of the activity of this gene may prove useful in the treatment of endocrine/metabolically related diseases, such as obesity and diabetes.

Moderate to low levels of expression of this gene is also seen in a number of cancer cell lines derived from colon, lung, and ovarian cancer. Therefore, therapeutic modulation of this gene may be useful in the treatment of colon, lung and ovarian cancers.

Panel 1.3D Summary: Ag2078 Three experiments with same probe-primer sets are in excellent agreement. Highest expression of this gene is seen in fetal kidney (CTs=26-27.8), with lower expression in the adult lung. This pattern correlates to the expression seen in panel 1.5. Please see panel 1.5 for further discussion of this gene.

Panel 4D Summary: Ag2078 Highest expression of this gene is detected in thymus (CT=27.3). This gene or its protein product may thus play an important role in T cell development. Small molecule therapeutics, or antibody therapeutics designed against the protein encoded for by this gene could be utilized to modulate immune function (T cell development) and be important for organ transplant, AIDS treatment or post chemotherapy immune reconstitiution.

Moderate to low levels of expression of this gene is also seen in lupus kidney, resting and cytokine activated mucoepidermoid NCI-H292 cells and dermal fibroblasts. Therefore, therapeutic modulation of this gene may be useful in the treatment of chronic obstructive pulmonary disease, asthma, allergy, emphysema, lupus kidney and skin disorders, including psoriasis.

Panel 5D Summary: Ag2078 Highest expression of this gene is detected in uterus and adipose of diabetic patients on insulin (CT=30.9-31). In addition, moderate to low levels of expression of this gene is also seen in uterus and placenta. Therefore, therapeutic modulation of this gene may be useful in the treatment of obesity and diabetes.

C. CG118051-02: ALDH8 splice variant, submitted to study DDSMT on 09/26/01 by saguo; classification type=Finished In-silico; novelty=Update-Variants; ORF start=407, ORF stop=1436, frame=2; 1586 bp.

Expression of gene CG118051-02 was assessed using the primer-probe set Ag3729, described in Table CA. Results of the RTQ-PCR runs are shown in Tables CB and CC.

Table CA. Probe Name Ag3729

Primers		Length	Start Position	SEQ ID No

360

Forward	5'-ttcaagaaaacaagcagcttct-3'	22	273	258
Probe	TET-5'-cccaggacctgcataagccagct-3'-TAMRA	23	309	259
Reverse	5'-ctcagatatgtctgcctcgaa-3'	21	332	260

Table CB. Panel 2.2

5

Tissue Name	Rel. Exp.(%) Ag3729, Run 174441818	Rel. Exp.(%) Ag3729, Run 259034396	Tissue Name	Rel. Exp.(%) Ag3729, Run 174441818	Rel. Exp.(%) Ag3729, Run 259034396
Normal Colon	0.4	0.3	Kidney Margin (OD04348)	0.0	0.0
Colon cancer (OD06064)	1.4	1.0	Kidney malignant cancer (OD06204B)	0.0	0.0
Colon Margin (OD06064)	0.0	0.0	Kidney normal adjacent tissue (OD06204E)	0.0	0.0
Colon cancer (OD06159)	0.2	0.1	Kidney Cancer (OD04450-01)	0.0	0.0
Colon Margin (OD06159)	0.0	0.0	Kidney Margin (OD04450-03)	1.3	0.9
Colon cancer (OD06297-04)	0.0	0.0	Kidney Cancer 8120613	0.0	0.0
Colon Margin (OD06297-05)	0.0	0.0	Kidney Margin 8120614	0.0	0.0
CC Gr.2 ascend colon (ODO3921)	1.1	0.8	Kidney Cancer 9010320	0.5	0.3
CC Margin (ODO3921)	0.0	0.0	Kidney Margin 9010321	1.8	1.4
Colon cancer metastasis (OD06104)	0.2	0.1	Kidney Cancer 8120607	0.0	0.0
Lung Margin (OD06104)	0.0	0.0	Kidney Margin 8120608	1.0	0.8
Colon mets to lung (OD04451-01)	0.2	0.2	Normal Uterus	0.0	0.0
Lung Margin (OD04451-02)	0.0	0.0	Uterine Cancer 064011	1.8	1.2
Normal Prostate	2.3	1.8	Normal Thyroid	0.0	0.0
Prostate Cancer (OD04410)	2.2	1.6	Thyroid Cancer 064010	0.0	0.0
Prostate Margin (OD04410)	5.1	3.8	Thyroid Cancer A302152	0.0	0.0

Normal Ovary	0.7	0.3	Thyroid Margin A302153	0.0	0.0
Ovarian cancer (OD06283-03)	2.5	1.7	Normal Breast	9.2	6.5
Ovarian Margin (OD06283-07)	0.0	0.0	Breast Cancer (OD04566)	17.4	12.9
Ovarian Cancer 064008	1.0	0.6	Breast Cancer 1024	**100.0**	**100.0**
Ovarian cancer (OD06145)	0.4	0.3	Breast Cancer (OD04590-01)	3.9	2.5
Ovarian Margin (OD06145)	0.5	0.3	Breast Cancer Mets (OD04590-03)	1.2	0.9
Ovarian cancer (OD06455-03)	0.9	0.5	Breast Cancer Metastasis (OD04655-05)	48.6	34.4
Ovarian Margin (OD06455-07)	0.0	0.0	Breast Cancer 064006	2.4	2.1
Normal Lung	0.0	0.0	Breast Cancer 9100266	55.1	43.8
Invasive poor diff. lung adeno (ODO4945-01	9.2	7.5	Breast Margin 9100265	14.7	10.8
Lung Margin (ODO4945-03)	0.0	0.0	Breast Cancer A209073	32.1	24.5
Lung Malignant Cancer (OD03126)	0.5	0.4	Breast Margin A2090734	9.1	6.4
Lung Margin (OD03126)	0.4	0.3	Breast cancer (OD06083)	69.7	61.6
Lung Cancer (OD05014A)	0.0	0.0	Breast cancer node metastasis (OD06083)	28.5	23.3
Lung Margin (OD05014B)	0.8	0.6	Normal Liver	0.0	0.0
Lung cancer (OD06081)	44.8	0.3	Liver Cancer 1026	0.0	0.0
Lung Margin (OD06081)	0.0	0.0	Liver Cancer 1025	0.8	0.6
Lung Cancer (OD04237-01)	3.1	2.6	Liver Cancer 6004-T	0.2	0.1
Lung Margin (OD04237-02)	0.4	0.3	Liver Tissue 6004-N	0.4	0.3
Ocular Melanoma Metastasis	0.0	0.0	Liver Cancer 6005-T	0.0	0.0
Ocular Melanoma Margin (Liver)	0.0	0.0	Liver Tissue 6005-N	0.0	0.0
Melanoma Metastasis	0.0	0.0	Liver Cancer 064003	0.0	0.0
Melanoma Margin (Lung)	0.3	0.2	Normal Bladder	0.0	0.0

Normal Kidney	0.0	0.0	Bladder Cancer 1023	3.2	2.3
Kidney Ca, Nuclear grade 2 (OD04338)	1.5	1.2	Bladder Cancer A302173	4.5	3.2
Kidney Margin (OD04338)	0.4	0.3	Normal Stomach	0.0	0.0
Kidney Ca Nuclear grade 1/2 (OD04339)	0.0	0.0	Gastric Cancer 9060397	0.5	0.3
Kidney Margin (OD04339)	0.0	0.0	Stomach Margin 9060396	2.1	1.4
Kidney Ca, Clear cell type (OD04340)	0.0	0.0	Gastric Cancer 9060395	2.5	1.7
Kidney Margin (OD04340)	0.4	0.3	Stomach Margin 9060394	1.8	1.1
Kidney Ca, Nuclear grade 3 (OD04348)	0.0	0.0	Gastric Cancer 064005	0.0	0.0

Table CC. Panel 4.1D

5

Tissue Name	Rel. Ep.(%) Ag3729, Run 170222887	Tissue Name	Rel. Exp.(%) Ag3729, Run 170222887
Secondary Th1 act	0.0	HUVEC IL-1beta	0.0
Secondary Th2 act	0.0	HUVEC IFN gamma	0.0
Secondary Tr1 act	0.0	HUVEC TNF alpha + IFN gamma	0.0
Secondary Th1 rest	0.0	HUVEC TNF alpha + IL4	0.0
Secondary Th2 rest	0.0	HUVEC IL-11	0.0
Secondary Tr1 rest	0.0	Lung Microvascular EC none	0.0
Primary Th1 act	0.0	Lung Microvascular EC TNFalpha + IL-1beta	0.0
Primary Th2 act	0.0	Microvascular Dermal EC none	0.0
Primary Tr1 act	0.0	Microsvasular Dermal EC TNFalpha + IL-1beta	0.0
Primary Th1 rest	0.0	Bronchial epithelium TNFalpha + ILbeta	26.8
Primary Th2 rest	0.0	Small airway epithelium none	25.5
Primary Tr1 rest	0.0	Small airway epithelium TNFalpha + IL-1beta	46.7
CD45RA CD4 lymphocyte act	0.0	Coronery artery SMC rest	0.0
CD45RO CD4 lymphocyte act	0.0	Coronery artery SMC TNFalpha + IL-1beta	0.0
CD8 lymphocyte act	0.0	Astrocytes rest	0.0
Secondary CD8 lymphocyte rest	0.0	Astrocytes TNFalpha + IL-1beta	0.0

Secondary CD8 lymphocyte act	0.0	KU-812 (Basophil) rest	0.0
CD4 lymphocyte none	0.0	KU-812 (Basophil) PMA/ionomycin	0.0
2ry Th1/Th2/Tr1_anti-CD95 CH11	0.0	CCD1106 (Keratinocytes) none	0.0
LAK cells rest	0.0	CCD1106 (Keratinocytes) TNFalpha + IL-1beta	6.7
LAK cells IL-2	0.0	Liver cirrhosis	0.0
LAK cells IL-2+IL-12	0.0	NCI-H292 none	100.0
LAK cells IL-2+IFN gamma	0.0	NCI-H292 IL-4	55.9
LAK cells IL-2+ IL-18	0.0	NCI-H292 IL-9	82.9
LAK cells PMA/ionomycin	0.0	NCI-H292 IL-13	58.2
NK Cells IL-2 rest	0.0	NCI-H292 IFN gamma	60.3
Two Way MLR 3 day	0.0	HPAEC none	0.0
Two Way MLR 5 day	0.0	HPAEC TNF alpha + IL-1 beta	0.0
Two Way MLR 7 day	0.0	Lung fibroblast none	0.0
PBMC rest	0.0	Lung fibroblast TNF alpha + IL-1 beta	0.0
PBMC PWM	0.0	Lung fibroblast IL-4	0.0
PBMC PHA-L	0.0	Lung fibroblast IL-9	0.0
Ramos (B cell) none	7.4	Lung fibroblast IL-13	0.0
Ramos (B cell) ionomycin	3.1	Lung fibroblast IFN gamma	0.0
B lymphocytes PWM	0.0	Dermal fibroblast CCD1070 rest	0.0
B lymphocytes CD40L and IL-4	0.0	Dermal fibroblast CCD1070 TNF alpha	0.0
EOL-1 dbcAMP	0.0	Dermal fibroblast CCD1070 IL-1 beta	0.0
EOL-1 dbcAMP PMA/ionomycin	0.0	Dermal fibroblast IFN gamma	0.0
Dendritic cells none	0.0	Dermal fibroblast IL-4	0.0
Dendritic cells LPS	0.0	Dermal Fibroblasts rest	0.0
Dendritic cells anti-CD40	0.0	Neutrophils TNFa+LPS	0.0
Monocytes rest	0.0	Neutrophils rest	0.0
Monocytes LPS	0.0	Colon	0.0
Macrophages rest	0.0	Lung	6.3
Macrophages LPS	0.0	Thymus	7.8
HUVEC none	0.0	Kidney	2.6
HUVEC starved	0.0		

CNS_neurodegeneration_v1.0 Summary: Ag3729 Expression of this gene is low/undetectable in all samples on this panel (CTs>35).

5   **Panel 2.2 Summary:** Ag3729 Two experiments with same probe-primer sets are in good agreement. Highest expression of this gene is seen in breast cancer (CTs=27-29).

Thus, expression of this gene could be used to differentiate between the breast cancer

samples and other samples on this panel.

In addition, moderate expression of this gene is also seen in cancer samples derived

from colon, breast, ovarian, lung, bladder, kidney and uterine cancers. Interestingly,

5   expression of gene higher cancer compared to the corresponding normal adjacent tissue.

Thus, expression of this gene may be used as diagnostic marker to detect the presence of

colon, breast, ovarian, lung, bladder, kidney and uterine cancers and also, therapeutic

modulation of the expression or function of this gene may be effective in the treatment of

these cancers.

10   **Panel 4.1D Summary:** Ag3729 Expression of this gene is restricted to a few

samples, with highest expression is seen in untreated NCI-H292 cells (CT=31.4). The gene

is also expressed in a cluster of treated and untreated samples derived from the NCI-H292

cell line, a human airway epithelial cell line that produces mucins. Mucus overproduction is

an important feature of bronchial asthma and chronic obstructive pulmonary disease

15   samples. Interestingly, the transcript is also expressed at lower but still significant levels in

small airway and bronchial epithelium treated with IL-1 beta and TNF-alpha and untreated

small airway epithelium. The expression of the transcript in this mucoepidermoid cell line

that is often used as a model for airway epithelium (NCI-H292 cells) suggests that this

transcript may be important in the proliferation or activation of airway epithelium.

20   Therefore, therapeutics designed with the protein encoded by the transcript may reduce or

eliminate symptoms caused by inflammation in lung epithelia in chronic obstructive

pulmonary disease, asthma, allergy, and emphysema.

D. CG140468-02: SERINE/THREONINE-PROTEIN KINASE PAK 1.

25   Expression of gene CG140468-02 was assessed using the primer-probe set Ag7054,

described in Table DA. Results of the RTQ-PCR runs are shown in Table DB. Please note

that CG140468-02 represents a full-length physical clone.

Table DA. Probe Name Ag7054

30

Primers	Sequence	Length	Start Position	SEQ ID No
Forward	5'-ggtttgagaagattgccaagc-3'	21	819	261

Probe	TET-5'-cctcactccactgattgctgcagcta-3'-TAMRA	27	850	262
Reverse	5'-ctgggggtgagtgtggtttttag-3'	21	898	263

Table DB. General screening panel v1.6

Tissue Name	Rel. Exp.(%) Ag7054, Run 282273878	issue Name	Rel. Exp.(%) Ag7054, Run 282273878
Adipose	3.6	Renal ca. TK-10	10.7
Melanoma* Hs688(A).T	7.3	Bladder	9.0
Melanoma* Hs688(B).T	6.6	Gastric ca. (liver met.) NCI-N87	30.6
Melanoma* M14	13.3	Gastric ca. KATO III	49.3
Melanoma* LOXIMVI	21.6	Colon ca. SW-948	7.8
Melanoma* SK-MEL-5	8.1	Colon ca. SW480	2.5
Squamous cell carcinoma SCC-4	7.7	Colon ca.* (SW480 met) SW620	11.8
Testis Pool	5.6	Colon ca. HT29	22.2
Prostate ca.* (bone met) PC-3	3.3	Colon ca. HCT-116	19.1
Prostate Pool	8.0	Colon ca. CaCo-2	34.6
Placenta	9.5	Colon cancer tissue	9.0
Uterus Pool	2.4	Colon ca. SW1116	4.5
Ovarian ca. OVCAR-3	100.0	Colon ca. Colo-205	10.2
Ovarian ca. SK-OV-3	16.4	Colon ca. SW-48	8.0
Ovarian ca. OVCAR-4	3.3	Colon Pool	9.1
Ovarian ca. OVCAR-5	35.1	Small Intestine Pool	8.9
Ovarian ca. IGROV-1	5.3	Stomach Pool	5.1
Ovarian ca. OVCAR-8	8.4	Bone Marrow Pool	3.4
Ovary	5.1	Fetal Heart	1.5
Breast ca. MCF-7	2.2	Heart Pool	3.7
Breast ca. MDA-MB-231	11.8	Lymph Node Pool	8.3
Breast ca. BT 549	4.2	Fetal Skeletal Muscle	8.1
Breast ca. T47D	7.7	Skeletal Muscle Pool	4.3
Breast ca. MDA-N	5.8	Spleen Pool	5.1
Breast Pool	8.8	Thymus Pool	7.6
Trachea	7.7	CNS cancer (glio/astro) U87-MG	6.3
Lung	4.1	CNS cancer (glio/astro) U-118-MG	12.7
Fetal Lung	7.9	CNS cancer (neuro;met) SK-N-AS	6.2
Lung ca. NCI-N417	7.9	CNS cancer (astro) SF-539	7.4
Lung ca. LX-1	19.9	CNS cancer (astro) SNB-75	14.1
Lung ca. NCI-H146	3.5	CNS cancer (glio) SNB-19	5.5
Lung ca. SHP-77	5.8	CNS cancer (glio) SF-295	5.8

Lung ca. A549	8.8	Brain (Amygdala) Pool	24.8
Lung ca. NCI-H526	3.5	Brain (cerebellum)	85.9
Lung ca. NCI-H23	11.0	Brain (fetal)	16.4
Lung ca. NCI-H460	1.0	Brain (Hippocampus) Pool	21.2
Lung ca. HOP-62	3.5	Cerebral Cortex Pool	64.6
Lung ca. NCI-H522	20.7	Brain (Substantia nigra) Pool	27.9
Liver	0.7	Brain (Thalamus) Pool	51.8
Fetal Liver	9.1	Brain (whole)	55.5
Liver ca. HepG2	0.5	Spinal Cord Pool	5.0
Kidney Pool	11.3	Adrenal Gland	4.9
Fetal Kidney	16.0	Pituitary gland Pool	4.9
Renal ca. 786-0	9.9	Salivary Gland	2.7
Renal ca. A498	4.4	Thyroid (female)	5.8
Renal ca. ACHN	6.9	Pancreatic ca. CAPAN2	9.7
Renal ca. UO-31	13.5	Pancreas Pool	5.5

General_screening_panel_v1.6 Summary: Ag7054 Highest expression of this gene is detected in a ovarian cancer cell line (CT=25.4). Moderate levels of expression of this gene is also seen in cluster of cancer cell lines derived from pancreatic, gastric, colon, lung, liver, renal, breast, ovarian, prostate, squamous cell carcinoma, melanoma and brain cancers. Thus, expression of this gene could be used as a marker to detect the presence of these cancers. Furthermore, therapeutic modulation of the expression or function of this gene may be effective in the treatment of pancreatic, gastric, colon, lung, liver, renal, breast, ovarian, prostate, squamous cell carcinoma, melanoma and brain cancers.

Among tissues with metabolic or endocrine function, this gene is expressed at moderate levels in pancreas, adipose, adrenal gland, thyroid, pituitary gland, skeletal muscle, heart, liver and the gastrointestinal tract. Therefore, therapeutic modulation of the activity of this gene may prove useful in the treatment of endocrine/metabolically related diseases, such as obesity and diabetes.

In addition, this gene is expressed at high levels in all regions of the central nervous system examined, including amygdala, hippocampus, substantia nigra, thalamus, cerebellum, cerebral cortex, and spinal cord. Therefore, therapeutic modulation of this gene product may be useful in the treatment of central nervous system disorders such as Alzheimer's disease, Parkinson's disease, epilepsy, multiple sclerosis, schizophrenia and depression.

Interestingly, this gene is expressed at much higher levels in fetal (CT=28.9) when compared to adult liver (CT=32.7). This observation suggests that expression of this gene can be used to distinguish fetal from adult liver. In addition, the relative overexpression of this gene in fetal tissue suggests that the protein product may enhance liver growth or

5   development in the fetus and thus may also act in a regenerative capacity in the adult. Therefore, therapeutic modulation of the protein encoded by this gene could be useful in treatment of liver related diseases.

E. CG142564-01: CARNITINE O-PALMITOYLTRANSFERASE I.

10   Expression of gene CG142564-01 was assessed using the primer-probe set Ag6952, described in Table EA. Results of the RTQ-PCR runs are shown in Table EB. Please note that CG142564-02 represents a full-length physical clone.

Table EA. Probe Name Ag6952

15

Primers		Length	Start Position	SEQ ID No
Forward	5'-tctgctaccaatcccagatcc-3'	21	434	264
Probe	TET-5'-tcgacccagagcagcacccca-3'-TAMRA	21	461	265
Reverse	5'-catctgctacagggccaaag-3'	20	504	266

Table EB. General screening panel v1.6

20

Tissue Name	Rel. Exp.(%) Ag6952, Run 278388893	Tissue Name	Rel. Exp.(%) Ag6952, Run 278388893
Adipose	4.1	Renal ca. TK-10	20.0
Melanoma* Hs688(A).T	0.8	Bladder	33.4
Melanoma* Hs688(B).T	1.2	Gastric ca. (liver met.) NCI-N87	81.2
Melanoma* M14	21.8	Gastric ca. KATO III	8.2
Melanoma* LOXIMVI	4.6	Colon ca. SW-948	5.4
Melanoma* SK-MEL-5	8.5	Colon ca. SW480	14.8
Squamous cell carcinoma SCC-4	1.6	Colon ca.* (SW480 met) SW620	17.1
Testis Pool	31.6	Colon ca. HT29	1.3
Prostate ca.* (bone met) PC-3	9.3	Colon ca. HCT-116	14.3

Prostate Pool	5.8	Colon ca. CaCo-2	6.7
Placenta	8.5	Colon cancer tissue	7.6
Uterus Pool	0.7	Colon ca. SW1116	4.4
Ovarian ca. OVCAR-3	5.0	Colon ca. Colo-205	4.7
Ovarian ca. SK-OV-3	50.7	Colon ca. SW-48	2.6
Ovarian ca. OVCAR-4	1.9	Colon Pool	3.4
Ovarian ca. OVCAR-5	25.3	Small Intestine Pool	2.9
Ovarian ca. IGROV-1	6.9	Stomach Pool	2.9
Ovarian ca. OVCAR-8	4.7	Bone Marrow Pool	1.5
Ovary	3.0	Fetal Heart	100.0
Breast ca. MCF-7	9.7	Heart Pool	42.6
Breast ca. MDA-MB-231	9.1	Lymph Node Pool	2.9
Breast ca. BT 549	14.3	Fetal Skeletal Muscle	17.9
Breast ca. T47D	3.3	Skeletal Muscle Pool	21.8
Breast ca. MDA-N	0.8	Spleen Pool	10.4
Breast Pool	3.1	Thymus Pool	17.9
Trachea	3.8	CNS cancer (glio/astro) U87-MG	12.3
Lung	3.0	CNS cancer (glio/astro) U-118-MG	25.3
Fetal Lung	7.3	CNS cancer (neuro;met) SK-N-AS	21.0
Lung ca. NCI-N417	1.2	CNS cancer (astro) SF-539	2.6
Lung ca. LX-1	22.8	CNS cancer (astro) SNB-75	16.5
Lung ca. NCI-H146	3.6	CNS cancer (glio) SNB-19	10.1
Lung ca. SHP-77	26.4	CNS cancer (glio) SF-295	61.1
Lung ca. A549	13.4	Brain (Amygdala) Pool	4.5
Lung ca. NCI-H526	0.8	Brain (cerebellum)	39.0
Lung ca. NCI-H23	13.8	Brain (fetal)	13.2
Lung ca. NCI-H460	13.9	Brain (Hippocampus) Pool	3.6
Lung ca. HOP-62	32.8	Cerebral Cortex Pool	3.4
Lung ca. NCI-H522	21.6	Brain (Substantia nigra) Pool	5.3
Liver	0.4	Brain (Thalamus) Pool	5.6
Fetal Liver	2.2	Brain (whole)	3.3
Liver ca. HepG2	5.0	Spinal Cord Pool	4.8
Kidney Pool	2.7	Adrenal Gland	6.9
Fetal Kidney	4.6	Pituitary gland Pool	3.2
Renal ca. 786-0	14.6	Salivary Gland	4.9
Renal ca. A498	1.8	Thyroid (female)	1.1
Renal ca. ACHN	7.6	Pancreatic ca. CAPAN2	12.1
Renal ca. UO-31	11.9	Pancreas Pool	5.0

General_screening_panel_v1.6 Summary: Ag6952 Highest expression of this gene is detected in fetal heart (CT=26.7). Moderate to high levels of expression of this gene

5   is also seen in tissues with metabolic/endocrine functions such as pancreas, adipose,

adrenal gland, thyroid, pituitary gland, skeletal muscle, heart, liver and the gastrointestinal

tract. Therefore, therapeutic modulation of the activity of this gene may prove useful in the

treatment of endocrine/metabolically related diseases, such as obesity and diabetes.

Moderate levels of expression of this gene is also seen in cluster of cancer cell lines

5   derived from pancreatic, gastric, colon, lung, liver, renal, breast, ovarian, prostate,

squamous cell carcinoma, melanoma and brain cancers. Thus, expression of this gene could

be used as a marker to detect the presence of these cancers. Furthermore, therapeutic

modulation of the expression or function of this gene may be effective in the treatment of

pancreatic, gastric, colon, lung, liver, renal, breast, ovarian, prostate, squamous cell

10   carcinoma, melanoma and brain cancers.

In addition, this gene is expressed at moderate levels in all regions of the central

nervous system examined, including amygdala, hippocampus, substantia nigra, thalamus,

cerebellum, cerebral cortex, and spinal cord. Therefore, therapeutic modulation of this gene

product may be useful in the treatment of central nervous system disorders such as

15   Alzheimer's disease, Parkinson's disease, epilepsy, multiple sclerosis, schizophrenia and

depression.

F. CG142797-01: Cathepsin L like.

Expression of gene CG142797-01 was assessed using the primer-probe set Ag7539,
described in Table FA.

20   **Table FA. Probe Name Ag7539**

Primers	Sequencs	Length	Start Position	SEQ ID No
Forward	5'-ctctaacacgtgaccacagtctaga-3'	25	68	267
Probe	TET-5'-tcttgtgctttgccttccacttggt-3'-TAMRA	25	103	268
Reverse	5'-atcttcatgttctccatgtcatataatc-3'	28	128	269

25   **CNS_neurodegeneration_v1.0 Summary:** Ag7539 Expression of this gene is

low/undetectable (CTs > 35) across all of the samples on this panel.

Panel 4.1D Summary: Ag7539 Expression of this gene is low/undetectable (CTs

> 35) across all of the samples on this panel.

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G. CG143216-01: Diacylglycerol Kinase.

Expression of gene CG143216-01 was assessed using the primer-probe sets Ag4554 and Ag7230, described in Tables GA and GB. Results of the RTQ-PCR runs are shown in Tables GC, GD, GE and GF.

Table GA. Probe Name Ag4554

Primers	Sequence	Length	Start Position	SEQ ID No
Forward	5'-aatgctccaggttcaattttct-3'	22	1349	270
Probe	TET-5'-accaaccagcaggaccagtttgactt-3'-TAMRA	26	1390	271
Reverse	5'-gacgcgataaacttcaacaaaa-3'	22	1419	272

Table GB. Probe Name Ag7230

Primers	Sequence	Length	Start Position	SEQ ID No
Forward	5'-gcatatcgttgttgggggact-3'	20	852	273
Probe	TET-5'-atggatgtgtcctcagtccaccacaa-3'-TAMRA	26	880	274
Reverse	5'-cacggagtagcgaaggagtg-3'	20	911	275

Table GC. CNS_neurodegeneration_v1.0

Tissue Name	Rel. Exp.(%) Ag4554, Run 224721290	Rel. Exp.(%) Ag7230, Run 288742189	issue Name	Rel. Exp.(%) Ag4554, Run 224721290	Rel. Exp.(%) Ag7230, Run 288742189
AD 1 Hippo	9.3	14.1	Control (Path) 3 Temporal Ctx	5.7	5.3
AD 2 Hippo	22.2	20.2	Control (Path) 4 Temporal Ctx	20.0	19.2
AD 3 Hippo	10.6	9.7	AD 1 Occipital Ctx	7.3	18.6
AD 4 Hippo	7.1	5.3	AD 2 Occipital Ctx (Missing)	0.0	0.0
AD 5 hippo	100.0	100.0	AD 3 Occipital Ctx	11.3	8.0
AD 6 Hippo	36.9	42.0	AD 4 Occipital Ctx	19.8	13.4
Control 2 Hippo	22.7	23.8	AD 5 Occipital Ctx	15.9	18.0
Control 4 Hippo	7.7	10.2	AD 6 Occipital Ctx	53.2	54.3

Control (Path) 3 Hippo	6.9	5.2	Control 1 Occipital Ctx	4.5	3.9
AD 1 Temporal Ctx	15.7	18.2	Control 2 Occipital Ctx	81.8	90.8
AD 2 Temporal Ctx	20.2	20.0	Control 3 Occipital Ctx	14.4	14.7
AD 3 Temporal Ctx	9.9	8.0	Control 4 Occipital Ctx	6.4	6.8
AD 4 Temporal Ctx	18.8	9.8	Control (Path) 1 Occipital Ctx	45.4	57.8
AD 5 Inf Temporal Ctx	97.9	81.2	Control (Path) 2 Occipital Ctx	6.1	6.1
AD 5 SupTemporal Ctx	31.6	36.3	Control (Path) 3 Occipital Ctx	5.1	5.2
AD 6 Inf Temporal Ctx	26.2	28.9	Control (Path) 4 Occipital Ctx	12.6	12.8
AD 6 Sup Temporal Ctx	29.1	33.7	Control 1 Parietal Ctx	6.4	5.7
Control 1 Temporal Ctx	9.5	5.1	Control 2 Parietal Ctx	26.4	26.4
Control 2 Temporal Ctx	39.0	43.2	Control 3 Parietal Ctx	18.0	19.6
Control 3 Temporal Ctx	10.1	11.4	Control (Path) 1 Parietal Ctx	56.3	70.7
Control 4 Temporal Ctx	6.6	6.7	Control (Path) 2 Parietal Ctx	15.7	15.2
Control (Path) 1 Temporal Ctx	32.8	35.1	Control (Path) 3 Parietal Ctx	5.5	5.1
Control (Path) 2 Temporal Ctx	20.4	22.8	Control (Path) 4 Parietal Ctx	41.5	36.3

Table GD. General screening panel v1.4

5

Tissue Name	Rel. Exp.(%) Ag4554, Run 222809973	issue Name	Rel. Exp.(%) Ag4554, Run 222809973
Adipose	5.4	Renal ca. TK-10	34.6
Melanoma* Hs688(A).T	45.1	Bladder	15.8
Melanoma* Hs688(B).T	45.1	Gastric ca. (liver met.) NCI-N87	21.3
Melanoma* M14	85.9	Gastric ca. KATO III	84.1
Melanoma* LOXIMVI	21.9	Colon ca. SW-948	0.7
Melanoma* SK-MEL-5	69.7	Colon ca. SW480	52.5
Squamous cell carcinoma SCC-4	26.8	Colon ca.* (SW480 met) SW620	27.0
Testis Pool	6.8	Colon ca. HT29	12.5

Prostate ca.* (bone met) PC-3	29.9	Colon ca. HCT-116	72.7
Prostate Pool	6.9	Colon ca. CaCo-2	25.5
Placenta	5.7	Colon cancer tissue	24.1
Uterus Pool	4.8	Colon ca. SW1116	8.5
Ovarian ca. OVCAR-3	14.9	Colon ca. Colo-205	12.9
Ovarian ca. SK-OV-3	**100.0**	Colon ca. SW-48	6.5
Ovarian ca. OVCAR-4	10.2	Colon Pool	15.0
Ovarian ca. OVCAR-5	36.1	Small Intestine Pool	17.8
Ovarian ca. IGROV-1	20.3	Stomach Pool	9.0
Ovarian ca. OVCAR-8	16.0	Bone Marrow Pool	5.0
Ovary	15.0	Fetal Heart	23.5
Breast ca. MCF-7	16.5	Heart Pool	12.2
Breast ca. MDA-MB-231	51.1	Lymph Node Pool	15.1
Breast ca. BT 549	47.3	Fetal Skeletal Muscle	4.6
Breast ca. T47D	62.0	Skeletal Muscle Pool	12.0
Breast ca. MDA-N	17.8	Spleen Pool	10.7
Breast Pool	12.5	Thymus Pool	26.2
Trachea	12.3	CNS cancer (glio/astro) U87-MG	65.1
Lung	1.2	CNS cancer (glio/astro) U-118-MG	79.0
Fetal Lung	27.4	CNS cancer (neuro;met) SK-N-AS	48.6
Lung ca. NCI-N417	8.0	CNS cancer (astro) SF-539	23.3
Lung ca. LX-1	52.1	CNS cancer (astro) SNB-75	89.5
Lung ca. NCI-H146	22.5	CNS cancer (glio) SNB-19	21.8
Lung ca. SHP-77	97.9	CNS cancer (glio) SF-295	63.7
Lung ca. A549	25.0	Brain (Amygdala) Pool	14.8
Lung ca. NCI-H526	0.0	Brain (cerebellum)	90.8
Lung ca. NCI-H23	45.1	Brain (fetal)	30.4
Lung ca. NCI-H460	15.9	Brain (Hippocampus) Pool	15.0
Lung ca. HOP-62	27.4	Cerebral Cortex Pool	29.3
Lung ca. NCI-H522	27.9	Brain (Substantia nigra) Pool	31.2
Liver	3.7	Brain (Thalamus) Pool	27.7
Fetal Liver	12.0	Brain (whole)	29.3
Liver ca. HepG2	28.1	Spinal Cord Pool	11.8
Kidney Pool	25.0	Adrenal Gland	29.1
Fetal Kidney	13.7	Pituitary gland Pool	24.8
Renal ca. 786-0	24.0	Salivary Gland	11.6
Renal ca. A498	4.5	Thyroid (female)	11.5
Renal ca. ACHN	6.3	Pancreatic ca. CAPAN2	10.4
Renal ca. UO-31	18.8	Pancreas Pool	21.8

Table GE. Panel 4.1D

Tissue Name	Rel. Exp.(%) Ag4554, Run 199319739	Rel. Exp.(%) Ag7230, Run 288211134	Tissue Name	Rel. Exp.(%) Ag4554, Run 199319739	Rel. Exp.(%) Ag7230, Run 288211134
Secondary Th1 act	70.2	48.3	HUVEC IL-1beta	62.9	38.4
Secondary Th2 act	44.8	30.4	HUVEC IFN gamma	50.3	35.1
Secondary Tr1 act	64.2	17.8	HUVEC TNF alpha + IFN gamma	18.2	14.0
Secondary Th1 rest	17.7	6.7	HUVEC TNF alpha + IL4	43.2	13.1
Secondary Th2 rest	22.4	6.6	HUVEC IL-11	38.2	16.7
Secondary Tr1 rest	17.0	6.0	Lung Microvascular EC none	100.0	100.0
Primary Th1 act	27.7	6.0	Lung Microvascular EC TNFalpha + IL-1beta	82.4	42.0
Primary Th2 act	42.3	24.8	Microvascular Dermal EC none	40.3	9.7
Primary Tr1 act	39.5	31.4	Microsvasular Dermal EC TNFalpha + IL-1beta	28.3	7.1
Primary Th1 rest	17.2	12.2	Bronchial epithelium TNFalpha + IL1beta	17.7	5.6
Primary Th2 rest	11.0	10.1	Small airway epithelium none	4.5	3.6
Primary Tr1 rest	39.2	1.2	Small airway epithelium TNFalpha + IL-1beta	11.4	6.6
CD45RA CD4 lymphocyte act	39.8	18.7	Coronery artery SMC rest	24.8	14.1
CD45RO CD4 lymphocyte act	44.4	31.4	Coronery artery SMC TNFalpha + IL-1beta	24.7	19.8
CD8 lymphocyte act	41.2	10.8	Astrocytes rest	11.7	10.2
Secondary CD8 lymphocyte rest	43.5	9.9	Astrocytes TNFalpha + IL-1beta	7.8	3.8
Secondary CD8 lymphocyte act	11.2	4.4	KU-812 (Basophil) rest	5.8	4.3
CD4 lymphocyte none	19.2	5.0	KU-812 (Basophil) PMA/ionomycin	7.9	5.7
2ry Th1/Th2/Tr1_anti-CD95 CH11	40.9	11.2	CCD1106 (Keratinocytes) none	14.6	13.4
LAK cells rest	21.0	8.0	CCD1106 (Keratinocytes) TNFalpha + IL-1beta	5.7	2.1

LAK cells IL-2	23.0	13.0	Liver cirrhosis	3.0	4.0
LAK cells IL-2+IL-12	12.7	1.5	NCI-H292 none	3.4	7.5
LAK cells IL-2+IFN gamma	14.6	5.6	NCI-H292 IL-4	7.1	8.0
LAK cells IL-2+ IL-18	18.7	7.7	NCI-H292 IL-9	9.7	6.6
LAK cells PMA/ionomycin	23.8	14.3	NCI-H292 IL-13	10.7	6.3
NK Cells IL-2 rest	42.9	35.8	NCI-H292 IFN gamma	3.2	1.5
Two Way MLR 3 day	22.5	9.9	HPAEC none	31.0	13.9
Two Way MLR 5 day	20.9	3.3	HPAEC TNF alpha + IL-1 beta	52.5	31.9
Two Way MLR 7 day	21.2	10.2	Lung fibroblast none	16.0	7.7
PBMC rest	12.0	6.8	Lung fibroblast TNF alpha + IL-1 beta	16.8	9.6
PBMC PWM	19.3	5.1	Lung fibroblast IL-4	16.3	7.6
PBMC PHA-L	29.9	14.4	Lung fibroblast IL-9	23.2	11.4
Ramos (B cell) none	19.3	6.5	Lung fibroblast IL-13	13.8	7.0
Ramos (B cell) ionomycin	21.3	13.7	Lung fibroblast IFN gamma	7.1	6.1
B lymphocytes PWM	18.2	9.9	Dermal fibroblast CCD1070 rest	22.7	36.6
B lymphocytes CD40L and IL-4	26.4	25.7	Dermal fibroblast CCD1070 TNF alpha	63.7	59.5
EOL-1 dbcAMP	29.3	26.2	Dermal fibroblast CCD1070 IL-1 beta	29.9	19.3
EOL-1 dbcAMP PMA/ionomycin	23.0	7.5	Dermal fibroblast IFN gamma	7.0	5.6
Dendritic cells none	28.9	17.6	Dermal fibroblast IL-4	20.6	12.9
Dendritic cells LPS	9.0	2.8	Dermal Fibroblasts rest	15.2	20.7
Dendritic cells anti-CD40	40.6	8.3	Neutrophils TNFa+LPS	18.4	16.0
Monocytes rest	20.7	7.6	Neutrophils rest	16.3	20.6
Monocytes LPS	18.2	15.7	Colon	14.1	3.9
Macrophages rest	20.0	8.2	Lung	9.9	2.6
Macrophages LPS	4.0	2.0	Thymus	39.2	7.4
HUVEC none	57.8	31.9	Kidney	18.8	11.6
HUVEC starved	64.2	50.0			

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Table GF. Panel 5 Islet

Tissue Name	Rel. Exp.() Ag4554, Run 306350410	Tissue Name	Rel. Exp.(%) Ag4554, Run 306350410
97457_Patient-02go_adipose	5.0	94709_Donor 2 AM - A_adipose	20.3
97476_Patient-07sk_skeletal muscle	0.0	94710_Donor 2 AM - B_adipose	12.8
97477_Patient-07ut_uterus	5.4	94711_Donor 2 AM - C_adipose	9.5
97478_Patient-07pl_placenta	2.6	94712_Donor 2 AD - A_adipose	18.0
99167_Bayer Patient 1	**100.0**	94713_Donor 2 AD - B_adipose	34.4
97482_Patient-08ut_uterus	2.4	94714_Donor 2 AD - C_adipose	17.3
97483_Patient-08pl_placenta	1.9	94742_Donor 3 U - A_Mesenchymal Stem Cells	10.0
97486_Patient-09sk_skeletal muscle	3.4	94743_Donor 3 U - B_Mesenchymal Stem Cells	9.7
97487_Patient-09ut_uterus	3.4	94730_Donor 3 AM - A_adipose	29.1
97488_Patient-09pl_placenta	0.9	94731_Donor 3 AM - B_adipose	47.0
97492_Patient-10ut_uterus	5.6	94732_Donor 3 AM - C_adipose	33.9
97493_Patient-10pl_placenta	6.0	94733_Donor 3 AD - A_adipose	46.3
97495_Patient-11go_adipose	4.7	94734_Donor 3 AD - B_adipose	72.7
97496_Patient-11sk_skeletal muscle	3.4	94735_Donor 3 AD - C_adipose	13.7
97497_Patient-11ut_uterus	6.0	77138_Liver_HepG2untreated	41.5
97498_Patient-11pl_placenta	2.0	73556_Heart_Cardiac stromal cells (primary)	8.5
97500_Patient-12go_adipose	8.7	81735_Small Intestine	18.0
97501_Patient-12sk_skeletal muscle	14.2	72409_Kidney_Proximal Convoluted Tubule	9.3
97502_Patient-12ut_uterus	12.3	82685_Small intestine_Duodenum	20.2
97503_Patient-12pl_placenta	3.5	90650_Adrenal_Adrenocortical adenoma	10.1
94721_Donor 2 U - A_Mesenchymal Stem Cells	21.6	72410_Kidney_HRCE	16.8
94722_Donor 2 U - B_Mesenchymal Stem Cells	6.3	72411_Kidney_HRE	6.8
94723_Donor 2 U - C_Mesenchymal Stem Cells	20.2	73139_Uterus_Uterine smooth muscle cells	19.5

5

CNS_neurodegeneration_v1.0 Summary: Ag4554/Ag7230 Two experiments with different probe-primer sets are in excellent agreement. This panel confirms the expression of this gene at low levels in the brains of an independent group of individuals.

However, no differential expression of this gene was detected between Alzheimer's diseased postmortem brains and those of non-demented controls in this experiment. Please see Panel 1.4 for a discussion of this gene in treatment of central nervous system disorders.

General_screening_panel_v1.4 Summary: Ag4554 Highest expression of this
5   gene is detected in a ovarian cancer cell line (CT=25.4). Moderate levels of expression of this gene is also seen in cluster of cancer cell lines derived from pancreatic, gastric, colon, lung, liver, renal, breast, ovarian, prostate, squamous cell carcinoma, melanoma and brain cancers. Thus, expression of this gene could be used as a marker to detect the presence of these cancers. Furthermore, therapeutic modulation of the expression or function of this
10 gene may be effective in the treatment of pancreatic, gastric, colon, lung, liver, renal, breast, ovarian, prostate, squamous cell carcinoma, melanoma and brain cancers.

Among tissues with metabolic or endocrine function, this gene is expressed at moderate levels in pancreas, adipose, adrenal gland, thyroid, pituitary gland, skeletal muscle, heart, liver and the gastrointestinal tract. Therefore, therapeutic modulation of the
15 activity of this gene may prove useful in the treatment of endocrine/metabolically related diseases, such as obesity and diabetes.

In addition, this gene is expressed at high levels in all regions of the central nervous system examined, including amygdala, hippocampus, substantia nigra, thalamus, cerebellum, cerebral cortex, and spinal cord. Therefore, therapeutic modulation of this gene
20 product may be useful in the treatment of central nervous system disorders such as Alzheimer's disease, Parkinson's disease, epilepsy, multiple sclerosis, schizophrenia and depression.

Interestingly, this gene is expressed at much higher levels in fetal (CT=27.3) when compared to adult lung (CT=31.8). This observation suggests that expression of this gene
25 can be used to distinguish fetal from adult lung. In addition, the relative overexpression of this gene in fetal tissue suggests that the protein product may enhance lung growth or development in the fetus and thus may also act in a regenerative capacity in the adult. Therefore, therapeutic modulation of the protein encoded by this gene could be useful in treatment of lung related diseases.

30 **Panel 4.1D Summary:** Ag4554/Ag7230 Two experiments with different probe-primer sets are in excellent agreement. Highest expression of this gene is detected in lung microvascular endothelial cells (CTs=28-29). This gene is expressed at high to moderate levels in a wide range of cell types of significance in the immune response in

health and disease. These cells include members of the T-cell, B-cell, endothelial cell, macrophage/monocyte, and peripheral blood mononuclear cell family, as well as epithelial and fibroblast cell types from lung and skin, and normal tissues represented by colon, lung, thymus and kidney. This ubiquitous pattern of expression suggests that this gene product may be involved in homeostatic processes for these and other cell types and tissues. This pattern is in agreement with the expression profile in General_screening_panel_v1.4 and also suggests a role for the gene product in cell survival and proliferation. Therefore, modulation of the gene product with a functional therapeutic may lead to the alteration of functions associated with these cell types and lead to improvement of the symptoms of patients suffering from autoimmune and inflammatory diseases such as asthma, allergies, inflammatory bowel disease, lupus erythematosus, psoriasis, rheumatoid arthritis, and osteoarthritis.

Panel 5 Islet Summary: Ag4554 Highest expression of this gene is detected in islet cells (CT=29.8). This gene shows a widespread expression pattern which correlates with the pattern seen in panel 1.4. Please see panel 1.4 for further discussion of this gene.

H. CG143787-01: Disintegrin Protease.

Expression of gene CG143787-01 was assessed using the primer-probe sets Ag6532, Ag6655 and Ag7048, described in Tables HA, HB and HC. Please note that CG143787-01 represents a full-length physical clone.

Table HA. Probe Name Ag6532

Primers	Sequencs	Length	Start Position	SEQ ID No
Forward	5'-atcatcaccaaagatacctttttatctc-3'	27	474	276
Probe	TET-5'-agaaaccaaagtgcctgctgcaagc-3'-TAMRA	25	501	277
Reverse	5'-gtgttgtcattatatttgtaggaataggt-3'	29	526	278

Table HB. Probe Name Ag6655

Primers	Sequenes	Length	Start Position	SEQ ID No
Forward	5'-atcatcaccaaagatacctttttatctc-3'	27	474	279

Probe	TET-5'-agaaaccaaagtgcctgctgcaagc-3'-TAMRA	25	501	280
Reverse	5'-gtgttgtcattatatttgtaggaataggt-3'	29	526	281

Table HC. Probe Name Ag7048

Primers	Sequence	Length	Start Position	SEQ ID No
Forward	5'-acatcatcaccaaagatacctttta-3'	25	472	282
Probe	TET-5'-caaagtgcctgctgcaagcacctatt-3'-TAMRA	26	507	283
Reverse	5'-gttcccacacactggtgttg-3'	20	549	284

General_screening_panel_v1.6 Summary: Ag6655/Ag7048 Expression of this gene is low/undetectable (CTs > 35) across all of the samples on this panel.

Panel 4.1D Summary: Ag6655 Expression of this gene is low/undetectable (CTs > 35) across all of the samples on this panel.

I. CG144112-01: NEUROPSIN PRECURSOR.

Expression of gene CG144112-01 was assessed using the primer-probe set Ag7123, described in Table IA. Please note that CG56663-01 represents a full-length physical clone.

Table IA. Probe Name Ag7123

Primers	Sequencs	Length	Start Position	SEQ ID No
Forward	5'-gcctgggcaggaaatacac-3'	19	353	285
Probe	TET-5'-tacgcctgggagaccacagcctacag-3'-TAMRA	26	325	286
Reverse	5'-tctcggggactgcacttct-3'	19	292	287

CNS_neurodegeneration_v1.0 Summary: Ag7123 Expression of this gene is low/undetectable (CTs > 35) across all of the samples on this panel.

Panel 4.1D Summary: Ag7123 Expression of this gene is low/undetectable (CTs > 35) across all of the samples on this panel.

J. CG144112-04: Kallikrein-8.

Expression of gene CG144112-04 was assessed using the primer-probe set Ag5271, described in Table JA.

Table JA. Probe Name Ag5271

5

Primers	Sequence	Length	Start Position	SEQ ID No
Forward	5'-gcagggcagggcgattct-3'	18	97	288
Probe	TET-5'-cacatcctggggctcagacccctgtg-3'-TAMRA	26	153	289
Reverse	5'-ctagaatcagcccttgctgccta-3'	23	245	290

CNS_neurodegeneration_v1.0 Summary: Ag5271 Expression of this gene is
10   low/undetectable (CTs > 35) across all of the samples on this panel.

Panel 4.1D Summary: Ag5271 Expression of this gene is low/undetectable (CTs > 35) across all of the samples on this panel.

K. CG144686-01: MAST CELL CARBOXYPEPTIDASE A PRECURSOR.

15   Expression of gene CG144686-01 was assessed using the primer-probe set Ag6864, described in Table KA. Results of the RTQ-PCR runs are shown in Tables KB and KC. Please note that CG144686-01 represents a full-length physical clone.

Table KA. Probe Name Ag6864

20

Primers	Sequencs	Length	Start Position	SEQ ID No
Forward	5'-aaccagtgagctccgaga-3'	18	122	291
Probe	TET-5'-caaatttggttttctccttccagaatcc-3'-TAMRA	28	146	292
Reverse	5'-tctgcacgttggctttat-3'	18	177	293

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Table KB. General screening panel v1.6

Tissue Name	Rel. Exp.(%) Ag6864, Run 278387547	issue Name	Rel. Exp.(%) Ag6864, Run 278387547
Adipose	15.0	Renal ca. TK-10	0.0
Melanoma* Hs688(A).T	0.3	Bladder	0.0
Melanoma* Hs688(B).T	0.7	Gastric ca. (liver met.) NCI-N87	0.0
Melanoma* M14	0.0	Gastric ca. KATO III	0.0
Melanoma* LOXIMVI	0.0	Colon ca. SW-948	0.0
Melanoma* SK-MEL-5	0.0	Colon ca. SW480	0.0
Squamous cell carcinoma SCC-4	0.0	Colon ca.* (SW480 met) SW620	0.0
Testis Pool	7.6	Colon ca. HT29	0.0
Prostate ca.* (bone met) PC-3	0.0	Colon ca. HCT-116	0.0
Prostate Pool	16.4	Colon ca. CaCo-2	0.0
Placenta	0.1	Colon cancer tissue	70.7
Uterus Pool	15.8	Colon ca. SW1116	0.0
Ovarian ca. OVCAR-3	0.0	Colon ca. Colo-205	0.0
Ovarian ca. SK-OV-3	0.0	Colon ca. SW-48	0.0
Ovarian ca. OVCAR-4	0.0	Colon Pool	78.5
Ovarian ca. OVCAR-5	0.0	Small Intestine Pool	0.0
Ovarian ca. IGROV-1	0.0	Stomach Pool	20.0
Ovarian ca. OVCAR-8	0.0	Bone Marrow Pool	23.2
Ovary	2.5	Fetal Heart	4.6
Breast ca. MCF-7	0.0	Heart Pool	20.0
Breast ca. MDA-MB-231	0.0	Lymph Node Pool	**100.0**
Breast ca. BT 549	0.7	Fetal Skeletal Muscle	5.5
Breast ca. T47D	0.0	Skeletal Muscle Pool	1.5
Breast ca. MDA-N	0.0	Spleen Pool	3.0
Breast Pool	0.0	Thymus Pool	18.2
Trachea	2.5	CNS cancer (glio/astro) U87-MG	0.0
Lung	2.7	CNS cancer (glio/astro) U-118-MG	1.8
Fetal Lung	5.3	CNS cancer (neuro;met) SK-N-AS	0.0
Lung ca. NCI-N417	0.0	CNS cancer (astro) SF-539	0.0
Lung ca. LX-1	0.0	CNS cancer (astro) SNB-75	0.0
Lung ca. NCI-H146	0.0	CNS cancer (glio) SNB-19	0.0
Lung ca. SHP-77	4.5	CNS cancer (glio) SF-295	0.0
Lung ca. A549	0.0	Brain (Amygdala) Pool	0.0
Lung ca. NCI-H526	0.0	Brain (cerebellum)	0.0
Lung ca. NCI-H23	0.0	Brain (fetal)	0.0
Lung ca. NCI-H460	0.0	Brain (Hippocampus) Pool	0.0
Lung ca. HOP-62	0.9	Cerebral Cortex Pool	0.0

Lung ca. NCI-H522	0.0	Brain (Substantia nigra) Pool	0.0
Liver	0.0	Brain (Thalamus) Pool	0.0
Fetal Liver	6.0	Brain (whole)	0.0
Liver ca. HepG2	0.0	Spinal Cord Pool	0.0
Kidney Pool	51.4	Adrenal Gland	0.7
Fetal Kidney	1.1	Pituitary gland Pool	1.0
Renal ca. 786-0	0.2	Salivary Gland	0.0
Renal ca. A498	0.0	Thyroid (female)	0.2
Renal ca. ACHN	0.0	Pancreatic ca. CAPAN2	0.0
Renal ca. UO-31	0.2	Pancreas Pool	10.4

Table KC. Panel 5 Islet

5

Tissue Name	Rel. Exp.(%) Ag6864 Run 305424858	Rel. Exp.(%) Ag6864, Run 307650498	Tissue Name	Rel. Exp.(%) Ag6864, Run 305424858	Rel. Exp.(%) Ag6864, Run 307650498
97457_Patient-02go_adipose	5.5	34.9	94709_Donor 2 AM - A_adipose	0.0	0.0
97476_Patient-07sk_skeletal muscle	0.0	0.0	94710_Donor 2 AM - B_adipose	0.0	0.0
97477_Patient-07ut_uterus	1.4	32.1	94711_Donor 2 AM - C_adipose	0.0	0.0
97478_Patient-07pl_placenta	0.0	4.7	94712_Donor 2 AD - A_adipose	0.0	0.0
99167_Bayer Patient 1	0.0	0.0	94713_Donor 2 AD - B_adipose	0.0	0.0
97482_Patient-08ut_uterus	0.0	0.0	94714_Donor 2 AD - C_adipose	2.3	0.0
97483_Patient-08pl_placenta	0.0	0.0	94742_Donor 3 U - A_Mesenchymal Stem Cells	0.0	0.0
97486_Patient-09sk_skeletal muscle	7.6	15.5	94743_Donor 3 U - B_Mesenchymal Stem Cells	0.0	0.0
97487_Patient-09ut_uterus	28.7	11.2	94730_Donor 3 AM - A_adipose	0.0	0.0
97488_Patient-09pl_placenta	1.4	0.0	94731_Donor 3 AM - B_adipose	0.0	1.9
97492_Patient-10ut_uterus	10.4	7.2	94732_Donor 3 AM - C_adipose	0.0	0.0
97493_Patient-10pl_placenta	0.0	5.9	94733_Donor 3 AD - A_adipose	0.0	0.0
97495_Patient-11go_adipose	20.0	5.0	94734_Donor 3 AD - B_adipose	0.0	0.0

97496_Patient-11sk_skeletal muscle	6.0	8.7	94735_Donor 3 AD - C_adipose	0.0	0.0
97497_Patient-11ut_uterus	45.1	65.1	77138_Liver_HepG2untreated	0.0	0.0
97498_Patient-11pl_placenta	0.0	0.0	73556_Heart_Cardiac stromal cells (primary)	5.1	3.2
97500_Patient-12go_adipose	59.9	59.9	81735_Small Intestine	73.2	65.1
97501_Patient-12sk_skeletal muscle	100.0	100.0	72409_Kidney_Proximal Convoluted Tubule	0.0	0.0
97502_Patient-12ut_uterus	29.1	97.3	82685_Small intestine_Duodenum	59.0	67.4
97503_Patient-12pl_placenta	5.0	2.3	90650_Adrenal_Adrenocortical adenoma	0.0	0.0
94721_Donor 2 U - A_Mesenchymal Stem Cells	0.0	0.0	72410_Kidney_HRCE	0.0	0.0
94722_Donor 2 U - B_Mesenchymal Stem Cells	0.0	0.0	72411_Kidney_HRE	0.0	0.0
94723_Donor 2 U - C_Mesenchymal Stem Cells	1.5	0.0	73139_Uterus_Uterine smooth muscle cells	0.0	0.0

General_screening_panel_v1.6 Summary: Ag6864 Highest expression of this gene is seen in lymph node (CT=29). Moderate levels of expression are also seen predominantly in normal tissue, including adipose, colon, heart, thymus, prostate, and kidney, as well as in colon cancer tissue. Thus, expression of this gene could be used to identify these samples and tissues. Modulation of the expression of this gene may also be effective in the treatment of diseases of these tissues, including cancer, obesity and diabetes.

Panel 5 Islet Summary: Ag6864 Two experiments with the same probe and primer produce results that are in excellent agreement. Highest expression of this gene is seen in skeletal muscle (CTs=33.5). Please see Panel 1.6 for discussion of this gene.

L. CG144906-01: TESTISIN PRECURSOR.

Expression of gene CG144906-01 was assessed using the primer-probe set Ag6915, described in Table LA. Please note that CG144906-01 represents a full-length physical clone.

Table LA. Probe Name Ag6915

Primers	Sequence	Length	Start Position	SEQ ID No
Forward	5'-catgccatcctccacattt-3'	19	337	294
Probe	TET-5'-cagcagtctgtccggttctcaaactc-3'-TAMRA	26	356	295
Reverse	5'-gtgcctcatcctctttgatgta-3'	22	398	296

General_screening_panel_v1.6 Summary: Ag6915 Expression of this gene is low/undetectable (CTs > 35) across all of the samples on this panel.

M. CG144997-01: RNase H I.

Expression of gene CG144997-01 was assessed using the primer-probe set Ag7057, described in Table MA. Results of the RTQ-PCR runs are shown in Table MB. Please note that CG144997-01 represents a full-length physical clone.

Table MA. Probe Name Ag7057

Primers		Length	Start Position	SEQ ID No
Forward	5'-gtaaacgccgattcctgct-3'	19	468	297
Probe	TET-5'-cttctacgcccattactggagcagca-3'-TAMRA	26	493	298
Reverse	5'-gaatgagtgcagagacacgttt-3'	22	558	299

Table MB. General_screening_panel_v1.6

Tissue Name	Rel. Exp.(%) Ag7057, Run 282273884	Issue Name	Rel. Exp.(%) Ag7057, Run 282273884
Adipose	3.9	Renal ca. TK-10	33.9
Melanoma* Hs688(A).T	23.8	Bladder	15.7
Melanoma* Hs688(B).T	28.3	Gastric ca. (liver met.) NCI-N87	49.0
Melanoma* M14	50.7	Gastric ca. KATO III	100.0
Melanoma* LOXIMVI	57.8	Colon ca. SW-948	11.4
Melanoma* SK-MEL-5	51.4	Colon ca. SW480	76.3
Squamous cell carcinoma SCC-4	22.5	Colon ca.* (SW480 met) SW620	34.9

Testis Pool	9.0	Colon ca. HT29	15.8
Prostate ca.* (bone met) PC-3	60.3	Colon ca. HCT-116	36.6
Prostate Pool	5.4	Colon ca. CaCo-2	42.0
Placenta	4.5	Colon cancer tissue	17.6
Uterus Pool	1.9	Colon ca. SW1116	5.4
Ovarian ca. OVCAR-3	31.2	Colon ca. Colo-205	10.4
Ovarian ca. SK-OV-3	31.4	Colon ca. SW-48	6.8
Ovarian ca. OVCAR-4	17.1	Colon Pool	9.5
Ovarian ca. OVCAR-5	39.0	Small Intestine Pool	5.7
Ovarian ca. IGROV-1	13.3	Stomach Pool	5.1
Ovarian ca. OVCAR-8	15.0	Bone Marrow Pool	3.3
Ovary	4.9	Fetal Heart	4.7
Breast ca. MCF-7	21.8	Heart Pool	4.5
Breast ca. MDA-MB-231	17.3	Lymph Node Pool	8.9
Breast ca. BT 549	24.8	Fetal Skeletal Muscle	4.0
Breast ca. T47D	9.5	Skeletal Muscle Pool	2.3
Breast ca. MDA-N	22.7	Spleen Pool	4.1
Breast Pool	12.3	Thymus Pool	8.2
Trachea	7.3	CNS cancer (glio/astro) U87-MG	55.5
Lung	1.9	CNS cancer (glio/astro) U-118-MG	49.7
Fetal Lung	8.6	CNS cancer (neuro;met) SK-N-AS	49.7
Lung ca. NCI-N417	10.1	CNS cancer (astro) SF-539	22.1
Lung ca. LX-1	22.4	CNS cancer (astro) SNB-75	45.1
Lung ca. NCI-H146	11.9	CNS cancer (glio) SNB-19	16.7
Lung ca. SHP-77	82.9	CNS cancer (glio) SF-295	56.6
Lung ca. A549	54.0	Brain (Amygdala) Pool	7.3
Lung ca. NCI-H526	8.9	Brain (cerebellum)	20.0
Lung ca. NCI-H23	37.9	Brain (fetal)	8.0
Lung ca. NCI-H460	37.1	Brain (Hippocampus) Pool	8.1
Lung ca. HOP-62	12.1	Cerebral Cortex Pool	12.0
Lung ca. NCI-H522	56.6	Brain (Substantia nigra) Pool	6.7
Liver	0.8	Brain (Thalamus) Pool	12.1
Fetal Liver	6.7	Brain (whole)	7.1
Liver ca. HepG2	18.6	Spinal Cord Pool	6.7
Kidney Pool	10.8	Adrenal Gland	6.9
Fetal Kidney	5.8	Pituitary gland Pool	2.9
Renal ca. 786-0	21.6	Salivary Gland	2.6
Renal ca. A498	17.1	Thyroid (female)	2.5
Renal ca. ACHN	17.6	Pancreatic ca. CAPAN2	23.3
Renal ca. UO-31	18.0	Pancreas Pool	6.0

General_screening_panel_v1.6 Summary: Ag7057 Highest expression of this gene is detected in a gastric cancer cell line (CT=27). Moderate levels of expression of this

gene is also seen in cluster of cancer cell lines derived from pancreatic, gastric, colon, lung, liver, renal, breast, ovarian, prostate, squamous cell carcinoma, melanoma and brain cancers. Thus, expression of this gene could be used as a marker to detect the presence of these cancers. Furthermore, therapeutic modulation of the expression or function of this

5   gene may be effective in the treatment of pancreatic, gastric, colon, lung, liver, renal, breast, ovarian, prostate, squamous cell carcinoma, melanoma and brain cancers.

Among tissues with metabolic or endocrine function, this gene is expressed at moderate levels in pancreas, adipose, adrenal gland, thyroid, pituitary gland, skeletal muscle, heart, liver and the gastrointestinal tract. Therefore, therapeutic modulation of the

10   activity of this gene may prove useful in the treatment of endocrine/metabolically related diseases, such as obesity and diabetes.

In addition, this gene is expressed at moderate levels in all regions of the central nervous system examined, including amygdala, hippocampus, substantia nigra, thalamus, cerebellum, cerebral cortex, and spinal cord. Therefore, therapeutic modulation of this gene

15   product may be useful in the treatment of central nervous system disorders such as Alzheimer's disease, Parkinson's disease, epilepsy, multiple sclerosis, schizophrenia and depression.

N. CG145494-01: PRESTIN.

Expression of gene CG145494-01 was assessed using the primer-probe sets

20   Ag6694, Ag7803 and Ag7797, described in Tables NA, NB and NC. Results of the RTQ-PCR runs are shown in Table ND.

Table NA. Probe Name Ag6694

Primers	Sequeces	Length	Start Position	SEQ ID No
Forward	5'-ggcacagaggccagagat-3'	18	559	300
Probe	TET-5'-gtgaccttactttcaggaatcattcagttttgc-3'-TAMRA	33	604	301
Reverse	5'-ggctctgtgagatatatggcc-3'	21	663	302

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Table NB. Probe Name Ag7803

Primers	Sequencs	Length	Start Position	SEQ ID No
Forward	5'-ggagaaccagcaaaatagagct-3'	22	1367	303
Probe	TET-5'-ccaatcccaggaacaaggaggacaca a-3'-TAMRA	27	1409	304
Reverse	5'-atcacagcagtgatcaaacca-3'	21	1440	305

5

Table NC. Probe Name Ag7797

Primers	Sequenes	Length	Start Position	SEQ ID No
Forward	5'-ccatctggcttaccacttttg-3'	21	1391	306
Probe	TET-5'-cacagcagtgatcaaaccatagtccaa tcc-3'-TAMRA	30	1429	307
Reverse	5'-aaatcacagtcagcagagcaat-3'	22	1462	308

10

Table ND. General_screening_panel_v1.6

Tissue Name	Rel. Exp.(%) Ag6694, Run 277223811	issue Name	Rel. Exp.(%) Ag6694, Run 277223811
Adipose	0.0	Renal ca. TK-10	0.0
Melanoma* Hs688(A).T	0.0	Bladder	0.0
Melanoma* Hs688(B).T	0.0	Gastric ca. (liver met.) NCI-N87	0.0
Melanoma* M14	0.0	Gastric ca. KATO III	0.0
Melanoma* LOXIMVI	0.0	Colon ca. SW-948	0.0
Melanoma* SK-MEL-5	0.0	Colon ca. SW480	0.0
Squamous cell carcinoma SCC-4	0.0	Colon ca.* (SW480 met) SW620	0.0
Testis Pool	0.0	Colon ca. HT29	0.0
Prostate ca.* (bone met) PC-3	100.0	Colon ca. HCT-116	0.0
Prostate Pool	0.9	Colon ca. CaCo-2	0.0
Placenta	0.0	Colon cancer tissue	0.0
Uterus Pool	0.0	Colon ca. SW1116	0.0
Ovarian ca. OVCAR-3	0.0	Colon ca. Colo-205	0.0
Ovarian ca. SK-OV-3	0.0	Colon ca. SW-48	0.0
Ovarian ca. OVCAR-4	0.0	Colon Pool	0.0
Ovarian ca. OVCAR-5	0.0	Small Intestine Pool	0.0

Ovarian ca. IGROV-1	0.0	Stomach Pool	0.0
Ovarian ca. OVCAR-8	0.0	Bone Marrow Pool	0.0
Ovary	0.0	Fetal Heart	0.0
Breast ca. MCF-7	0.0	Heart Pool	0.0
Breast ca. MDA-MB-231	0.0	Lymph Node Pool	0.0
Breast ca. BT 549	0.0	Fetal Skeletal Muscle	0.0
Breast ca. T47D	0.0	Skeletal Muscle Pool	0.0
Breast ca. MDA-N	0.0	Spleen Pool	0.0
Breast Pool	0.0	Thymus Pool	0.0
Trachea	1.0	CNS cancer (glio/astro) U87-MG	0.0
Lung	0.0	CNS cancer (glio/astro) U-118-MG	0.0
Fetal Lung	2.9	CNS cancer (neuro;met) SK-N-AS	0.0
Lung ca. NCI-N417	0.0	CNS cancer (astro) SF-539	0.0
Lung ca. LX-1	0.0	CNS cancer (astro) SNB-75	0.0
Lung ca. NCI-H146	0.0	CNS cancer (glio) SNB-19	0.0
Lung ca. SHP-77	0.0	CNS cancer (glio) SF-295	0.0
Lung ca. A549	0.0	Brain (Amygdala) Pool	0.0
Lung ca. NCI-H526	0.0	Brain (cerebellum)	14.6
Lung ca. NCI-H23	0.0	Brain (fetal)	0.0
Lung ca. NCI-H460	0.0	Brain (Hippocampus) Pool	0.0
Lung ca. HOP-62	0.0	Cerebral Cortex Pool	0.0
Lung ca. NCI-H522	0.0	Brain (Substantia nigra) Pool	0.0
Liver	0.0	Brain (Thalamus) Pool	0.0
Fetal Liver	0.0	Brain (whole)	0.0
Liver ca. HepG2	0.0	Spinal Cord Pool	0.0
Kidney Pool	0.0	Adrenal Gland	0.0
Fetal Kidney	0.0	Pituitary gland Pool	0.0
Renal ca. 786-0	0.0	Salivary Gland	0.0
Renal ca. A498	0.0	Thyroid (female)	0.0
Renal ca. ACHN	0.0	Pancreatic ca. CAPAN2	0.0
Renal ca. UO-31	0.0	Pancreas Pool	0.0

CNS_neurodegeneration_v1.0 Summary: Ag7797 Expression of this gene is low/undetectable (CTs > 34.7) across all of the samples on this panel.

General_screening_panel_v1.6 Summary: Ag6694 Moderate level of expression of this gene is restricted to prostate cancer cell line (CT=32.6). Therefore, expression of this gene may be used to distinguish this sample from other samples in this panel and also as diagnostic marker to detect the presence of prostate cancer. In addition, therapeutic modulation of this gene may be useful in the treatment of prostate cancer.

Panel 4.1D Summary: Ag7803 Expression of this gene is low/undetectable (CTs > 35) across all of the samples on this panel.

O. CG145722-01: WEE1-like protein kinase.

Expression of gene CG145722-01 was assessed using the primer-probe set Ag6231, described in Table OA. Results of the RTQ-PCR runs are shown in Table OB.

Table OA. Probe Name Ag6231

Primers	Sequence	Length	Start Position	SEQ ID No
Forward	5'-gcttcctggctaatgagatttt-3'	22	1339	309
Probe	TET-5'-agaggattaccggcaccttcccaaag-3'-TAMRA	26	1364	310
Reverse	5'-tgttaatcccaaggcaaatatg-3'	22	1394	311

Table OB. General screening panel v1.5

Tissue Name	Rel. Exp.(%) Ag6231, Run 259211049	issue Name	Rel. Exp.(%) Ag6231, Run 259211049
Adipose	0.0	Renal ca. TK-10	0.0
Melanoma* Hs688(A).T	0.0	Bladder	0.0
Melanoma* Hs688(B).T	0.0	Gastric ca. (liver met.) NCI-N87	0.0
Melanoma* M14	0.0	Gastric ca. KATO III	0.0
Melanoma* LOXIMVI	0.0	Colon ca. SW-948	0.0
Melanoma* SK-MEL-5	0.0	Colon ca. SW480	0.0
Squamous cell carcinoma SCC-4	0.0	Colon ca.* (SW480 met) SW620	0.0
Testis Pool	0.0	Colon ca. HT29	0.0
Prostate ca.* (bone met) PC-3	0.0	Colon ca. HCT-116	0.0
Prostate Pool	0.0	Colon ca. CaCo-2	97.3
Placenta	0.0	Colon cancer tissue	0.0
Uterus Pool	0.0	Colon ca. SW1116	0.0
Ovarian ca. OVCAR-3	0.0	Colon ca. Colo-205	0.0
Ovarian ca. SK-OV-3	0.0	Colon ca. SW-48	0.0
Ovarian ca. OVCAR-4	0.0	Colon Pool	0.0
Ovarian ca. OVCAR-5	0.0	Small Intestine Pool	0.0
Ovarian ca. IGROV-1	0.0	Stomach Pool	0.0
Ovarian ca. OVCAR-8	0.0	Bone Marrow Pool	0.0

Ovary	0.0	Fetal Heart	0.0
Breast ca. MCF-7	0.0	Heart Pool	0.0
Breast ca. MDA-MB-231	0.0	Lymph Node Pool	0.0
Breast ca. BT 549	0.0	Fetal Skeletal Muscle	0.0
Breast ca. T47D	0.0	Skeletal Muscle Pool	0.0
Breast ca. MDA-N	0.0	Spleen Pool	0.0
Breast Pool	0.0	Thymus Pool	0.0
Trachea	0.0	CNS cancer (glio/astro) U87-MG	0.0
Lung	0.0	CNS cancer (glio/astro) U-118-MG	0.0
Fetal Lung	0.0	CNS cancer (neuro;met) SK-N-AS	0.0
Lung ca. NCI-N417	0.0	CNS cancer (astro) SF-539	0.0
Lung ca. LX-1	0.0	CNS cancer (astro) SNB-75	4.2
Lung ca. NCI-H146	**100.0**	CNS cancer (glio) SNB-19	0.0
Lung ca. SHP-77	2.3	CNS cancer (glio) SF-295	0.0
Lung ca. A549	0.0	Brain (Amygdala) Pool	2.3
Lung ca. NCI-H526	0.0	Brain (cerebellum)	5.6
Lung ca. NCI-H23	0.0	Brain (fetal)	2.6
Lung ca. NCI-H460	0.0	Brain (Hippocampus) Pool	0.0
Lung ca. HOP-62	0.0	Cerebral Cortex Pool	0.0
Lung ca. NCI-H522	0.0	Brain (Substantia nigra) Pool	0.0
Liver	0.0	Brain (Thalamus) Pool	0.0
Fetal Liver	0.0	Brain (whole)	3.7
Liver ca. HepG2	0.0	Spinal Cord Pool	0.0
Kidney Pool	1.8	Adrenal Gland	0.0
Fetal Kidney	2.2	Pituitary gland Pool	0.0
Renal ca. 786-0	0.0	Salivary Gland	0.0
Renal ca. A498	0.0	Thyroid (female)	0.0
Renal ca. ACHN	0.0	Pancreatic ca. CAPAN2	4.6
Renal ca. UO-31	6.0	Pancreas Pool	0.0

CNS_neurodegeneration_v1.0 Summary: Ag6231 Expression of this gene is low/undetectable (CTs > 35) across all of the samples on this panel.

General_screening_panel_v1.5 Summary: Ag6231 Low levels of expression of this gene is restricted to a lung cancer and a colon cancer cell lines (CTs=32.2). Therefore, expression of this gene may be used to distinguish these cell lines from other samples in this panel and also as diagnostic marker to detect the presence of colon and lung cancers. In addition, therapeutic modulation of this gene may be useful in the treatment of these cancers.

Panel 4.1D Summary: Ag6231 Expression of this gene is low/undetectable (CTs > 35) across all of the samples on this panel.

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P. CG145754-02: KALLIKREIN 7 PRECURSOR: [illegible]

Expression of gene CG145754-02 was assessed using the primer-probe set Ag7038, described in Table PA. Results of the RTQ-PCR runs are shown in Tables PB and PC. Please note that CG145754-02 represents a full-length physical clone.

Table PA. Probe Name Ag7038

Primers	Sequence	Length	Start Position	SEQ ID No
Forward	5'-tgttaatgacctcaagctcatctc-3'	24	342	312
Probe	TET-5'-ccccaggactgcacgaaggtttacaa-3'-TAMRA	26	367	313
Reverse	5'-tttcttggagtcggggatg-3'	19	426	314

Table PB. General_screening_panel_v1.6

Tissue Name	Rel. Exp.(%) Ag7038, Run 282273672	issue Name	Rel. Exp.(%) Ag7038, Run 282273672
Adipose	1.6	Renal ca. TK-10	0.0
Melanoma* Hs688(A).T	0.0	Bladder	0.0
Melanoma* Hs688(B).T	0.0	Gastric ca. (liver met.) NCI-N87	100.0
Melanoma* M14	0.0	Gastric ca. KATO III	22.1
Melanoma* LOXIMVI	0.0	Colon ca. SW-948	4.4
Melanoma* SK-MEL-5	0.0	Colon ca. SW480	10.5
Squamous cell carcinoma SCC-4	3.0	Colon ca.* (SW480 met) SW620	0.0
Testis Pool	0.0	Colon ca. HT29	0.0
Prostate ca.* (bone met) PC-3	0.0	Colon ca. HCT-116	9.7
Prostate Pool	0.0	Colon ca. CaCo-2	0.0
Placenta	0.0	Colon cancer tissue	0.6
Uterus Pool	0.0	Colon ca. SW1116	38.7
Ovarian ca. OVCAR-3	4.1	Colon ca. Colo-205	0.0
Ovarian ca. SK-OV-3	0.0	Colon ca. SW-48	0.0
Ovarian ca. OVCAR-4	3.1	Colon Pool	0.0
Ovarian ca. OVCAR-5	0.0	Small Intestine Pool	0.0
Ovarian ca. IGROV-1	0.0	Stomach Pool	0.0
Ovarian ca. OVCAR-8	0.0	Bone Marrow Pool	0.0
Ovary	0.0	Fetal Heart	0.0
Breast ca. MCF-7	0.0	Heart Pool	0.0

Breast ca. MDA-MB-231	0.0	Lymph Node Pool	0.0
Breast ca. BT 549	0.0	Fetal Skeletal Muscle	0.0
Breast ca. T47D	0.0	Skeletal Muscle Pool	0.0
Breast ca. MDA-N	0.0	Spleen Pool	0.0
Breast Pool	0.0	Thymus Pool	0.0
Trachea	0.0	CNS cancer (glio/astro) U87-MG	0.0
Lung	0.0	CNS cancer (glio/astro) U-118-MG	0.0
Fetal Lung	0.0	CNS cancer (neuro;met) SK-N-AS	0.0
Lung ca. NCI-N417	0.0	CNS cancer (astro) SF-539	0.0
Lung ca. LX-1	0.5	CNS cancer (astro) SNB-75	2.0
Lung ca. NCI-H146	0.0	CNS cancer (glio) SNB-19	0.0
Lung ca. SHP-77	0.0	CNS cancer (glio) SF-295	0.0
Lung ca. A549	0.0	Brain (Amygdala) Pool	1.5
Lung ca. NCI-H526	0.0	Brain (cerebellum)	5.6
Lung ca. NCI-H23	4.2	Brain (fetal)	0.0
Lung ca. NCI-H460	0.0	Brain (Hippocampus) Pool	4.0
Lung ca. HOP-62	0.0	Cerebral Cortex Pool	3.1
Lung ca. NCI-H522	0.0	Brain (Substantia nigra) Pool	1.4
Liver	0.0	Brain (Thalamus) Pool	3.9
Fetal Liver	0.0	Brain (whole)	0.2
Liver ca. HepG2	0.0	Spinal Cord Pool	0.3
Kidney Pool	0.0	Adrenal Gland	0.0
Fetal Kidney	1.3	Pituitary gland Pool	0.0
Renal ca. 786-0	0.0	Salivary Gland	0.0
Renal ca. A498	0.6	Thyroid (female)	0.0
Renal ca. ACHN	0.0	Pancreatic ca. CAPAN2	2.2
Renal ca. UO-31	0.0	Pancreas Pool	0.0

Table PC. Panel 5 Islet

5

Tissue Name	Rel. Exp.(%) Ag703, Run 305424861	Tissue Name	Rel. Exp.(%) Ag7038, Run 305424861
97457_Patient-02go_adipose	3.0	94709_Donor 2 AM - A_adipose	0.0
97476_Patient-07sk_skeletal muscle	0.0	94710_Donor 2 AM - B_adipose	100.0
97477_Patient-07ut_uterus	0.0	94711_Donor 2 AM - C_adipose	0.0
97478_Patient-07pl_placenta	0.0	94712_Donor 2 AD - A_adipose	0.0
99167_Bayer Patient 1	0.0	94713_Donor 2 AD - B_adipose	0.0
97482_Patient-08ut_uterus	0.0	94714_Donor 2 AD - C_adipose	0.0

97483_Patient-08pl_placenta	0.0	94742_Donor 3 U - A_Mesenchymal Stem Cells	13.0
97486_Patient-09sk_skeletal muscle	0.0	94743_Donor 3 U - B_Mesenchymal Stem Cells	5.5
97487_Patient-09ut_uterus	0.0	94730_Donor 3 AM - A_adipose	0.0
97488_Patient-09pl_placenta	0.0	94731_Donor 3 AM - B_adipose	0.0
97492_Patient-10ut_uterus	0.0	94732_Donor 3 AM - C_adipose	0.0
97493_Patient-10pl_placenta	0.0	94733_Donor 3 AD - A_adipose	0.0
97495_Patient-11go_adipose	2.7	94734_Donor 3 AD - B_adipose	0.0
97496_Patient-11sk_skeletal muscle	0.0	94735_Donor 3 AD - C_adipose	0.0
97497_Patient-11ut_uterus	0.0	77138_Liver_HepG2untreated	0.0
97498_Patient-11pl_placenta	0.0	73556_Heart_Cardiac stromal cells (primary)	0.0
97500_Patient-12go_adipose	1.5	81735_Small Intestine	0.0
97501_Patient-12sk_skeletal muscle	0.0	72409_Kidney_Proximal Convoluted Tubule	2.4
97502_Patient-12ut_uterus	1.0	82685_Small intestine_Duodenum	0.0
97503_Patient-12pl_placenta	0.0	90650_Adrenal_Adrenocortical adenoma	0.0
94721_Donor 2 U - A_Mesenchymal Stem Cells	0.0	72410_Kidney_HRCE	5.7
94722_Donor 2 U - B_Mesenchymal Stem Cells	0.0	72411_Kidney_HRE	10.2
94723_Donor 2 U - C_Mesenchymal Stem Cells	0.0	73139_Uterus_Uterine smooth muscle cells	0.0

General_screening_panel_v1.6 Summary: Ag7038 Highest expression of this gene is detected in a gastric cancer NCI-N87 cell line (CT=31.3). Expression of this gene seems to be restricted to number of colon and gastric cancer cell lines. Therefore, expression of this gene may be used to distinguish colon and gastric cancer cell lines from other samples in this panel and also as a diagnostic marker to detect the presence of colon and gastric cancers. In addition, therapeutic modulation of this gene may be useful in the treatment of colon and gastric cancer.

Panel 5 Islet Summary: Ag7038 Low levels of expression of this gene is restricted to adipose tissue (CT=33). Therefore, expression of this gene may be used to distinguish this adipose sample from other samples in this panel. In addition, therapeutic modulation of this gene may be useful in the treatment of metabolic diseases such as obesity and diabetes.

Another experiment (Run 307650500) with this probe-primer set showed
low/undetectable (CTs > 35) across all of the samples on this panel.

Q. CG145754-03: Kallikrein-7.

Expression of gene CG145754-03 was assessed using the primer-probe set Ag5272,
5   described in Table QA. Results of the RTQ-PCR runs are shown in Table QB.

Table QA. Probe Name Ag5272

Primers		Length	Start Position	SEQ ID No
Forward	5'-ggcagccaggggtgacaa-3'	18	119	315
Probe	TET-5'-cgccccatgtgcaagaggctccc-3'-TAMRA	23	149	316
Reverse	5'-cctccgcagtggagctgatt-3'	20	201	317

10

Table QB. Panel 4.1D

Tissue Name	Rel. Ep.(%) Ag5272, Run 230500478	Tissue Name	Rel. Exp.(%) Ag5272, Run 230500478
Secondary Th1 act	0.0	HUVEC IL-1beta	0.0
Secondary Th2 act	0.0	HUVEC IFN gamma	0.0
Secondary Tr1 act	0.0	HUVEC TNF alpha + IFN gamma	0.0
Secondary Th1 rest	0.0	HUVEC TNF alpha + IL4	0.0
Secondary Th2 rest	0.0	HUVEC IL-11	0.0
Secondary Tr1 rest	0.0	Lung Microvascular EC none	0.0
Primary Th1 act	0.0	Lung Microvascular EC TNFalpha + IL-1beta	0.0
Primary Th2 act	0.0	Microvascular Dermal EC none	0.0
Primary Tr1 act	0.0	Microsvasular Dermal EC TNFalpha + IL-1beta	0.0
Primary Th1 rest	0.0	Bronchial epithelium TNFalpha + IL1beta	1.3
Primary Th2 rest	0.0	Small airway epithelium none	100.0
Primary Tr1 rest	0.0	Small airway epithelium TNFalpha + IL-1beta	46.7
CD45RA CD4 lymphocyte act	0.6	Coronery artery SMC rest	0.0
CD45RO CD4 lymphocyte act	0.0	Coronery artery SMC TNFalpha + IL-1beta	0.0

CD8 lymphocyte act	0.0	Astrocytes rest	0.0
Secondary CD8 lymphocyte rest	0.0	Astrocytes TNFalpha + IL-1beta	0.0
Secondary CD8 lymphocyte act	0.0	KU-812 (Basophil) rest	0.0
CD4 lymphocyte none	0.0	KU-812 (Basophil) PMA/ionomycin	1.2
2ry Th1/Th2/Tr1_anti-CD95 CH11	0.0	CCD1106 (Keratinocytes) none	14.2
LAK cells rest	0.0	CCD1106 (Keratinocytes) TNFalpha + IL-1beta	4.5
LAK cells IL-2	0.0	Liver cirrhosis	0.0
LAK cells IL-2+IL-12	0.0	NCI-H292 none	0.0
LAK cells IL-2+IFN gamma	0.0	NCI-H292 IL-4	0.0
LAK cells IL-2+ IL-18	0.0	NCI-H292 IL-9	0.0
LAK cells PMA/ionomycin	0.0	NCI-H292 IL-13	0.6
NK Cells IL-2 rest	0.0	NCI-H292 IFN gamma	0.0
Two Way MLR 3 day	0.0	HPAEC none	0.0
Two Way MLR 5 day	0.0	HPAEC TNF alpha + IL-1 beta	0.0
Two Way MLR 7 day	0.0	Lung fibroblast none	0.0
PBMC rest	0.0	Lung fibroblast TNF alpha + IL-1 beta	0.0
PBMC PWM	0.0	Lung fibroblast IL-4	0.0
PBMC PHA-L	0.0	Lung fibroblast IL-9	0.0
Ramos (B cell) none	0.0	Lung fibroblast IL-13	0.0
Ramos (B cell) ionomycin	0.0	Lung fibroblast IFN gamma	0.0
B lymphocytes PWM	0.0	Dermal fibroblast CCD1070 rest	0.0
B lymphocytes CD40L and IL-4	0.0	Dermal fibroblast CCD1070 TNF alpha	0.0
EOL-1 dbcAMP	0.0	Dermal fibroblast CCD1070 IL-1 beta	0.0
EOL-1 dbcAMP PMA/ionomycin	0.0	Dermal fibroblast IFN gamma	0.0
Dendritic cells none	0.0	Dermal fibroblast IL-4	0.0
Dendritic cells LPS	0.5	Dermal Fibroblasts rest	0.0
Dendritic cells anti-CD40	0.0	Neutrophils TNFa+LPS	0.0
Monocytes rest	0.0	Neutrophils rest	0.0
Monocytes LPS	0.0	Colon	0.0
Macrophages rest	0.0	Lung	0.0
Macrophages LPS	0.0	Thymus	0.0
HUVEC none	0.0	Kidney	11.2
HUVEC starved	0.0		

Panel 4.1D Summary: Ag5272 Highest expression of this gene is seen in resting small airway epithelium (CT=32). Significant expression of this gene is also seen in

5   cytokines TNF-a and IL-1b treated small airway epithelium. Therefore, modulation of the

expression or activity of the protein encoded by this transcript through the application of small molecule therapeutics may be useful in the treatment of asthma, COPD, and emphysema.

R. CG146279-01: Potassium channel subfamily K member 10.

5	Expression of gene CG146279-01 was assessed using the primer-probe set Ag6035, described in Table RA. Results of the RTQ-PCR runs are shown in Tables RB, RC, RD and RE.

Table RA. Probe Name Ag6035

10

Primers		Length	Start Position	SEQ ID No
Forward	5'-atgaaatttccaatcgagacg-3'	21	61	318
Probe	TET-5'-ctaaagtggccgttcccgcagc-3'-TAMRA	22	107	319
Reverse	5'-ggggttgcccgttagtg-3'	17	156	320

Table RB. CNS_neurodegeneration_v1.0

15

Tissue Name	Rel. Exp.(%) Ag6035, Run 225246892	issue Name	Rel. Exp.(%) Ag6035, Run 225246892
AD 1 Hippo	22.5	Control (Path) 3 Temporal Ctx	9.9
AD 2 Hippo	25.9	Control (Path) 4 Temporal Ctx	38.2
AD 3 Hippo	12.4	AD 1 Occipital Ctx	22.2
AD 4 Hippo	13.5	AD 2 Occipital Ctx (Missing)	0.0
AD 5 Hippo	82.9	AD 3 Occipital Ctx	5.3
AD 6 Hippo	74.2	AD 4 Occipital Ctx	35.4
Control 2 Hippo	21.5	AD 5 Occipital Ctx	40.9
Control 4 Hippo	19.3	AD 6 Occipital Ctx	17.7
Control (Path) 3 Hippo	8.2	Control 1 Occipital Ctx	4.8
AD 1 Temporal Ctx	24.3	Control 2 Occipital Ctx	53.2
AD 2 Temporal Ctx	43.8	Control 3 Occipital Ctx	39.2
AD 3 Temporal Ctx	4.5	Control 4 Occipital Ctx	8.2
AD 4 Temporal Ctx	36.6	Control (Path) 1 Occipital Ctx	88.3
AD 5 Inf Temporal Ctx	100.0	Control (Path) 2 Occipital Ctx	7.1
AD 5 Sup Temporal Ctx	62.0	Control (Path) 3 Occipital Ctx	2.5

AD 6 Inf Temporal Ctx	74.7	Control (Path) 4 Occipital Ctx	37.1
AD 6 Sup Temporal Ctx	65.1	Control 1 Parietal Ctx	8.9
Control 1 Temporal Ctx	5.8	Control 2 Parietal Ctx	77.4
Control 2 Temporal Ctx	29.5	Control 3 Parietal Ctx	17.1
Control 3 Temporal Ctx	22.7	Control (Path) 1 Parietal Ctx	77.9
Control 3 Temporal Ctx	22.7	Control (Path) 2 Parietal Ctx	22.4
Control (Path) 1 Temporal Ctx	74.2	Control (Path) 3 Parietal Ctx	6.3
Control (Path) 2 Temporal Ctx	47.0	Control (Path) 4 Parietal Ctx	51.4

Table RC. General screening panel v1.5

5

Tissue Name	Rel. Exp.(%) Ag6035, Run 228763481	issue Name	Rel. Exp.(%) Ag6035, Run 228763481
Adipose	0.5	Renal ca. TK-10	9.3
Melanoma* Hs688(A).T	0.0	Bladder	2.6
Melanoma* Hs688(B).T	0.0	Gastric ca. (liver met.) NCI-N87	8.2
Melanoma* M14	0.0	Gastric ca. KATO III	12.8
Melanoma* LOXIMVI	0.0	Colon ca. SW-948	1.0
Melanoma* SK-MEL-5	0.0	Colon ca. SW480	14.8
Squamous cell carcinoma SCC-4	0.0	Colon ca.* (SW480 met) SW620	29.1
Testis Pool	1.3	Colon ca. HT29	1.7
Prostate ca.* (bone met) PC-3	0.0	Colon ca. HCT-116	12.7
Prostate Pool	4.7	Colon ca. CaCo-2	12.3
Placenta	2.0	Colon cancer tissue	5.3
Uterus Pool	2.5	Colon ca. SW1116	0.0
Ovarian ca. OVCAR-3	3.3	Colon ca. Colo-205	3.7
Ovarian ca. SK-OV-3	2.8	Colon ca. SW-48	3.4
Ovarian ca. OVCAR-4	3.8	Colon Pool	0.9
Ovarian ca. OVCAR-5	7.0	Small Intestine Pool	1.5
Ovarian ca. IGROV-1	10.4	Stomach Pool	2.1
Ovarian ca. OVCAR-8	3.1	Bone Marrow Pool	0.5
Ovary	1.1	Fetal Heart	1.3
Breast ca. MCF-7	3.7	Heart Pool	0.2
Breast ca. MDA-MB-231	6.9	Lymph Node Pool	0.9
Breast ca. BT 549	2.0	Fetal Skeletal Muscle	1.4
Breast ca. T47D	1.1	Skeletal Muscle Pool	2.3
Breast ca. MDA-N	4.3	Spleen Pool	0.6
Breast Pool	4.9	Thymus Pool	2.8
Trachea	0.2	CNS cancer (glio/astro) U87-MG	0.0
Lung	1.1	CNS cancer (glio/astro) U-118-MG	2.8

Fetal Lung	4.1	CNS cancer (neuro;met) SK-N-AS	2.0
Lung ca. NCI-N417	3.9	CNS cancer (astro) SF-539	2.2
Lung ca. LX-1	30.1	CNS cancer (astro) SNB-75	4.6
Lung ca. NCI-H146	8.4	CNS cancer (glio) SNB-19	4.4
Lung ca. SHP-77	33.4	CNS cancer (glio) SF-295	11.4
Lung ca. A549	15.3	Brain (Amygdala) Pool	15.1
Lung ca. NCI-H526	4.8	Brain (cerebellum)	100.0
Lung ca. NCI-H23	5.1	Brain (fetal)	92.7
Lung ca. NCI-H460	7.9	Brain (Hippocampus) Pool	32.1
Lung ca. HOP-62	0.0	Cerebral Cortex Pool	21.8
Lung ca. NCI-H522	0.0	Brain (Substantia nigra) Pool	18.4
Liver	0.5	Brain (Thalamus) Pool	24.8
Fetal Liver	2.0	Brain (whole)	29.9
Liver ca. HepG2	7.4	Spinal Cord Pool	16.3
Kidney Pool	1.6	Adrenal Gland	2.2
Fetal Kidney	3.5	Pituitary gland Pool	3.7
Renal ca. 786-0	2.4	Salivary Gland	1.0
Renal ca. A498	2.4	Thyroid (female)	2.0
Renal ca. ACHN	11.8	Pancreatic ca. CAPAN2	0.0
Renal ca. UO-31	6.2	Pancreas Pool	0.6

Table RD. Panel 4.1D

5

Tissue Name	Rel. Exp.() Ag6035, Run 225157775	Tissue Name	Rel. Exp.(%) Ag6035, Run 225157775
Secondary Th1 act	0.0	HUVEC IL-1beta	0.0
Secondary Th2 act	0.0	HUVEC IFN gamma	0.0
Secondary Tr1 act	0.0	HUVEC TNF alpha + IFN gamma	0.0
Secondary Th1 rest	0.0	HUVEC TNF alpha + IL4	0.0
Secondary Th2 rest	0.0	HUVEC IL-11	0.0
Secondary Tr1 rest	0.0	Lung Microvascular EC none	0.0
Primary Th1 act	0.0	Lung Microvascular EC TNFalpha + IL-1beta	0.0
Primary Th2 act	0.0	Microvascular Dermal EC none	0.0
Primary Tr1 act	0.0	Microsvasular Dermal EC TNFalpha + IL-1beta	0.0
Primary Th1 rest	0.0	Bronchial epithelium TNFalpha + IL1beta	0.0
Primary Th2 rest	0.0	Small airway epithelium none	0.0
Primary Tr1 rest	0.0	Small airway epithelium TNFalpha + IL-1beta	0.0

CD45RA CD4 lymphocyte act	0.0	Coronery artery SMC rest	0.0
CD45RO CD4 lymphocyte act	0.0	Coronery artery SMC TNFalpha + IL-1beta	0.0
CD8 lymphocyte act	0.0	Astrocytes rest	0.0
Secondary CD8 lymphocyte rest	0.0	Astrocytes TNFalpha + IL-1beta	0.0
Secondary CD8 lymphocyte act	0.0	KU-812 (Basophil) rest	0.0
CD4 lymphocyte none	0.0	KU-812 (Basophil) PMA/ionomycin	0.0
2ry Th1/Th2/Tr1_anti-CD95 CH11	0.0	CCD1106 (Keratinocytes) none	0.0
LAK cells rest	0.0	CCD1106 (Keratinocytes) TNFalpha + IL-1beta	0.0
LAK cells IL-2	0.0	Liver cirrhosis	0.0
LAK cells IL-2+IL-12	0.0	NCI-H292 none	0.0
LAK cells IL-2+IFN gamma	0.0	NCI-H292 IL-4	0.0
LAK cells IL-2+ IL-18	0.0	NCI-H292 IL-9	0.0
LAK cells PMA/ionomycin	0.0	NCI-H292 IL-13	0.0
NK Cells IL-2 rest	0.0	NCI-H292 IFN gamma	0.0
Two Way MLR 3 day	10.1	HPAEC none	0.0
Two Way MLR 5 day	0.0	HPAEC TNF alpha + IL-1 beta	0.0
Two Way MLR 7 day	0.0	Lung fibroblast none	0.0
PBMC rest	5.5	Lung fibroblast TNF alpha + IL-1 beta	0.0
PBMC PWM	0.0	Lung fibroblast IL-4	0.0
PBMC PHA-L	0.0	Lung fibroblast IL-9	0.0
Ramos (B cell) none	0.0	Lung fibroblast IL-13	0.0
Ramos (B cell) ionomycin	0.0	Lung fibroblast IFN gamma	0.0
B lymphocytes PWM	0.0	Dermal fibroblast CCD1070 rest	0.0
B lymphocytes CD40L and IL-4	0.0	Dermal fibroblast CCD1070 TNF alpha	0.0
EOL-1 dbcAMP	**100.0**	Dermal fibroblast CCD1070 IL-1 beta	0.0
EOL-1 dbcAMP PMA/ionomycin	36.1	Dermal fibroblast IFN gamma	0.0
Dendritic cells none	0.0	Dermal fibroblast IL-4	0.0
Dendritic cells LPS	0.0	Dermal Fibroblasts rest	0.0
Dendritic cells anti-CD40	0.0	Neutrophils TNFa+LPS	0.0
Monocytes rest	9.9	Neutrophils rest	0.0
Monocytes LPS	0.0	Colon	0.0
Macrophages rest	0.0	Lung	0.0
Macrophages LPS	0.0	Thymus	8.5
HUVEC none	0.0	Kidney	7.7
HUVEC starved	0.0		

Table RE. Panel 5 Islet

Tissue Name	Rel. Exp.(%) Ag6035 Run 253578284	Rel. Exp.(%) Ag6035, Run 306414003	Tissue Name	Rel. Exp.(%) Ag6035, Run 253578284	Rel. Exp.(%) Ag6035, Run 306414003
97457_Patient-02go_adipose	0.0	0.0	94709_Donor 2 AM - A_adipose	0.0	0.0
97476_Patient-07sk_skeletal muscle	0.0	0.0	94710_Donor 2 AM - B_adipose	0.0	0.0
97477_Patient-07ut_uterus	0.0	0.0	94711_Donor 2 AM - C_adipose	0.0	0.0
97478_Patient-07pl_placenta	0.0	0.0	94712_Donor 2 AD - A_adipose	0.0	0.0
99167_Bayer Patient 1	100.0	100.0	94713_Donor 2 AD - B_adipose	0.0	0.0
97482_Patient-08ut_uterus	0.0	0.0	94714_Donor 2 AD - C_adipose	0.0	0.0
97483_Patient-08pl_placenta	0.0	0.0	94742_Donor 3 U - A_Mesenchymal Stem Cells	0.0	0.0
97486_Patient-09sk_skeletal muscle	0.0	0.0	94743_Donor 3 U - B_Mesenchymal Stem Cells	0.0	0.0
97487_Patient-09ut_uterus	0.0	0.0	94730_Donor 3 AM - A_adipose	0.0	0.0
97488_Patient-09pl_placenta	0.0	0.0	94731_Donor 3 AM - B_adipose	0.0	0.0
97492_Patient-10ut_uterus	0.0	0.0	94732_Donor 3 AM - C_adipose	0.0	0.0
97493_Patient-10pl_placenta	0.0	0.0	94733_Donor 3 AD - A_adipose	0.0	0.0
97495_Patient-11go_adipose	0.0	0.0	94734_Donor 3 AD - B_adipose	0.0	0.0
97496_Patient-11sk_skeletal muscle	0.0	0.0	94735_Donor 3 AD - C_adipose	0.0	0.0
97497_Patient-11ut_uterus	0.0	0.0	77138_Liver_HepG2untreated	0.0	0.0
97498_Patient-11pl_placenta	0.0	0.0	73556_Heart_Cardiac stromal cells (primary)	0.0	0.0
97500_Patient-12go_adipose	0.0	0.0	81735_Small Intestine	0.0	0.0
97501_Patient-12sk_skeletal muscle	0.0	0.0	72409_Kidney_Proximal Convoluted Tubule	0.0	0.0
97502_Patient-12ut_uterus	0.0	0.0	82685_Small intestine_Duodenum	0.0	0.0
97503_Patient-12pl_placenta	0.0	0.0	90650_Adrenal_Adrenocortical adenoma	0.0	16.2

94721_Donor 2 U - A_Mesenchymal Stem Cells	0.0	0.0	72410_Kidney_HRCE	0.0	0.0
94722_Donor 2 U - B_Mesenchymal Stem Cells	0.0	0.0	72411_Kidney_HRE	0.0	0.0
94723_Donor 2 U - C_Mesenchymal Stem Cells	0.0	0.0	73139_Uterus_Uterine smooth muscle cells	0.0	0.0

CNS_neurodegeneration_v1.0 Summary: Ag6035 This panel confirms the expression of this gene at low levels in the brains of an independent group of individuals. However, no differential expression of this gene was detected between Alzheimer's diseased postmortem brains and those of non-demented controls in this experiment. Please see Panel 1.5 for a discussion of this gene in treatment of central nervous system disorders.

General_screening_panel_v1.5 Summary: Ag6035 Highest expression of this gene is detected in cerebellum (CT=27). This gene codes for a splice variant of potassium channel TREK2. As reported in literature (Bang et al., 2000, J Biol Chem 275(23):17412-9, PMID: 10747911), this gene shows expression preferentially in all the regions of brain. Therefore, therapeutic modulation of this gene product may be useful in the treatment of central nervous system disorders such as Alzheimer's disease, Parkinson's disease, epilepsy, multiple sclerosis, schizophrenia and depression.

Moderate to low levels of expression of this gene is also seen in number of cancer cell lines derived from brain, colon, gastric, renal, lung, breast and ovarian cancer. Therefore, therapeutic modulation of this gene may be useful in the treatment of these cancers.

In addition, low levels of expression of this gene is also seen in tissues with metabolic/endocrine functions, including pancreas, adipose, adrenal gland, thyroid, pituitary gland, skeletal muscle, heart, liver and the gastrointestinal tract. Therefore, therapeutic modulation of the activity of this gene may prove useful in the treatment of endocrine/metabolically related diseases, such as obesity and diabetes.

Panel 4.1D Summary: Ag6035 Highest expression of this gene is detected in eosinophils (CT=32.5). Low levels of expression of this gene is also seen in PMA/ionomycin treated eosinophils. Therefore, therapeutic modulation of this gene or its protein product may useful in the treatment of hematopoietic disorders involving

eosinophils, parasitic infections, autoimmune and inflammatory diseases including allergy and asthma.

Panel 5 Islet Summary: Ag6035 Two experiments with same probe-primer sets are in excellent agreement. Low levels of expression of this gene are restricted to islet cells (CTs=33-34). This gene codes for a splice variant of potassium channel TREK2. Potassium channels play an important role in insulin secretion by islet beta cells upon stimulation by glucose. Alteration in the insulin secretion pathway through the use of sulfonylureas or genetic inactivation of K(ATP) channels may lead to inappropriate insulin secretion at low glucose (Henquin JC., 2000, Diabetes 49(11):1751-60, PMID: 11078440). Therefore, therapeutic modulation of this gene or its protein product may be useful in the treatment type 2 diabetes.

S. CG146403-01: Diacylglycerol acyltransferase 2.

Expression of gene CG146403-01 was assessed using the primer-probe set Ag6034, described in Table SA. Results of the RTQ-PCR runs are shown in Tables SB, SC and SD.

Table SA. Probe Name Ag6034

Primers	Sequence	Length	Start Position	SEQ ID No
Forward	5'-tggggagaatgacatctttaga-3'	22	540	321
Probe	TET-5'-cttaaggcttttgccacaggctcctg-3'-TAMRA	26	562	322
Reverse	5'-agagaagcccatgagcttctt-3'	21	613	323

Table SB. General screening panel v1.5

Tissue Name	Rel. Exp.(%) Ag6034, Run 228763480	issue Name	Rel. Exp.(%) Ag6034, Run 228763480
Adipose	0.2	Renal ca. TK-10	27.9
Melanoma* Hs688(A).T	0.0	Bladder	1.2
Melanoma* Hs688(B).T	0.0	Gastric ca. (liver met.) NCI-N87	0.5
Melanoma* M14	0.1	Gastric ca. KATO III	7.9
Melanoma* LOXIMVI	0.0	Colon ca. SW-948	3.6
Melanoma* SK-MEL-5	0.2	Colon ca. SW480	12.5

Squamous cell carcinoma SCC-4	0.0	Colon ca.* (SW480 met) SW620	1.9
Testis Pool	0.2	Colon ca. HT29	22.7
Prostate ca.* (bone met) PC-3	0.4	Colon ca. HCT-116	0.0
Prostate Pool	0.0	Colon ca. CaCo-2	**100.0**
Placenta	0.0	Colon cancer tissue	63.3
Uterus Pool	0.0	Colon ca. SW1116	0.0
Ovarian ca. OVCAR-3	0.0	Colon ca. Colo-205	2.1
Ovarian ca. SK-OV-3	0.2	Colon ca. SW-48	50.0
Ovarian ca. OVCAR-4	0.0	Colon Pool	0.2
Ovarian ca. OVCAR-5	0.1	Small Intestine Pool	0.4
Ovarian ca. IGROV-1	0.1	Stomach Pool	0.0
Ovarian ca. OVCAR-8	0.2	Bone Marrow Pool	0.0
Ovary	0.0	Fetal Heart	0.2
Breast ca. MCF-7	0.0	Heart Pool	0.1
Breast ca. MDA-MB-231	0.0	Lymph Node Pool	0.0
Breast ca. BT 549	0.1	Fetal Skeletal Muscle	0.1
Breast ca. T47D	0.0	Skeletal Muscle Pool	0.0
Breast ca. MDA-N	0.0	Spleen Pool	0.0
Breast Pool	0.0	Thymus Pool	0.1
Trachea	0.0	CNS cancer (glio/astro) U87-MG	0.0
Lung	0.0	CNS cancer (glio/astro) U-118-MG	0.0
Fetal Lung	0.4	CNS cancer (neuro;met) SK-N-AS	0.0
Lung ca. NCI-N417	0.1	CNS cancer (astro) SF-539	0.2
Lung ca. LX-1	28.1	CNS cancer (astro) SNB-75	0.0
Lung ca. NCI-H146	0.0	CNS cancer (glio) SNB-19	0.0
Lung ca. SHP-77	0.0	CNS cancer (glio) SF-295	0.0
Lung ca. A549	0.7	Brain (Amygdala) Pool	0.0
Lung ca. NCI-H526	0.0	Brain (cerebellum)	0.1
Lung ca. NCI-H23	0.0	Brain (fetal)	0.2
Lung ca. NCI-H460	4.2	Brain (Hippocampus) Pool	0.0
Lung ca. HOP-62	0.0	Cerebral Cortex Pool	0.1
Lung ca. NCI-H522	0.2	Brain (Substantia nigra) Pool	0.0
Liver	1.7	Brain (Thalamus) Pool	0.0
Fetal Liver	55.9	Brain (whole)	1.1
Liver ca. HepG2	62.9	Spinal Cord Pool	0.0
Kidney Pool	0.0	Adrenal Gland	0.0
Fetal Kidney	5.1	Pituitary gland Pool	0.0
Renal ca. 786-0	0.0	Salivary Gland	0.0
Renal ca. A498	0.1	Thyroid (female)	0.0
Renal ca. ACHN	0.0	Pancreatic ca. CAPAN2	0.0
Renal ca. UO-31	0.0	Pancreas Pool	0.0

Table SC. Panel 4.1D

Tissue Name	Rel. Ep.(%) Ag6034, Run 225245213	Tissue Name	Rel. Exp.(%) Ag6034, Run 225245213
Secondary Th1 act	0.0	HUVEC IL-1beta	0.0
Secondary Th2 act	0.0	HUVEC IFN gamma	0.0
Secondary Tr1 act	0.4	HUVEC TNF alpha + IFN gamma	0.0
Secondary Th1 rest	0.0	HUVEC TNF alpha + IL4	0.0
Secondary Th2 rest	0.0	HUVEC IL-11	0.0
Secondary Tr1 rest	0.0	Lung Microvascular EC none	0.0
Primary Th1 act	0.0	Lung Microvascular EC TNFalpha + IL-1beta	0.0
Primary Th2 act	0.0	Microvascular Dermal EC none	0.0
Primary Tr1 act	0.0	Microsvasular Dermal EC TNFalpha + IL-1beta	0.0
Primary Th1 rest	0.0	Bronchial epithelium TNFalpha + IL1beta	0.0
Primary Th2 rest	0.0	Small airway epithelium none	0.0
Primary Tr1 rest	0.0	Small airway epithelium TNFalpha + IL-1beta	0.0
CD45RA CD4 lymphocyte act	0.0	Coronery artery SMC rest	0.0
CD45RO CD4 lymphocyte act	0.0	Coronery artery SMC TNFalpha + IL-1beta	0.6
CD8 lymphocyte act	0.0	Astrocytes rest	0.0
Secondary CD8 lymphocyte rest	0.0	Astrocytes TNFalpha + IL-1beta	0.0
Secondary CD8 lymphocyte act	0.0	KU-812 (Basophil) rest	0.0
CD4 lymphocyte none	0.0	KU-812 (Basophil) PMA/ionomycin	0.0
2ry Th1/Th2/Tr1_anti-CD95 CH11	0.0	CCD1106 (Keratinocytes) none	0.0
LAK cells rest	0.0	CCD1106 (Keratinocytes) TNFalpha + IL-1beta	0.0
LAK cells IL-2	0.0	Liver cirrhosis	17.0
LAK cells IL-2+IL-12	0.0	NCI-H292 none	0.0
LAK cells IL-2+IFN gamma	0.0	NCI-H292 IL-4	0.0
LAK cells IL-2+ IL-18	0.0	NCI-H292 IL-9	0.0
LAK cells PMA/ionomycin	0.0	NCI-H292 IL-13	0.0
NK Cells IL-2 rest	0.0	NCI-H292 IFN gamma	0.0
Two Way MLR 3 day	0.0	HPAEC none	0.0
Two Way MLR 5 day	0.0	HPAEC TNF alpha + IL-1 beta	0.0
Two Way MLR 7 day	0.0	Lung fibroblast none	0.0
PBMC rest	0.0	Lung fibroblast TNF alpha + IL-1 beta	0.0

PBMC PWM	0.9	Lung fibroblast IL-4	0.0
PBMC PHA-L	0.0	Lung fibroblast IL-9	0.0
Ramos (B cell) none	0.0	Lung fibroblast IL-13	0.0
Ramos (B cell) ionomycin	0.0	Lung fibroblast IFN gamma	0.0
B lymphocytes PWM	0.0	Dermal fibroblast CCD1070 rest	0.0
B lymphocytes CD40L and IL-4	0.0	Dermal fibroblast CCD1070 TNF alpha	0.0
EOL-1 dbcAMP	0.0	Dermal fibroblast CCD1070 IL-1 beta	0.0
EOL-1 dbcAMP PMA/ionomycin	0.0	Dermal fibroblast IFN gamma	0.0
Dendritic cells none	0.0	Dermal fibroblast IL-4	0.0
Dendritic cells LPS	0.0	Dermal Fibroblasts rest	0.3
Dendritic cells anti-CD40	0.5	Neutrophils TNFa+LPS	4.0
Monocytes rest	0.0	Neutrophils rest	0.0
Monocytes LPS	0.0	Colon	81.2
Macrophages rest	0.0	Lung	4.7
Macrophages LPS	0.0	Thymus	18.0
HUVEC none	0.0	Kidney	**100.0**
HUVEC starved	0.0		

Table SD. Panel 5 Islet

Tissue Name	Rel. Exp.(%) Ag603, Run 256791126	Tissue Name	Rel. Exp.(%) Ag6034, Run 256791126
97457_Patient-02go_adipose	0.0	94709_Donor 2 AM - A_adipose	0.0
97476_Patient-07sk_skeletal muscle	0.0	94710_Donor 2 AM - B_adipose	0.0
97477_Patient-07ut_uterus	0.0	94711_Donor 2 AM - C_adipose	0.0
97478_Patient-07pl_placenta	0.0	94712_Donor 2 AD - A_adipose	0.0
99167_Bayer Patient 1	0.0	94713_Donor 2 AD - B_adipose	0.0
97482_Patient-08ut_uterus	0.0	94714_Donor 2 AD - C_adipose	0.0
97483_Patient-08pl_placenta	0.0	94742_Donor 3 U - A_Mesenchymal Stem Cells	0.0
97486_Patient-09sk_skeletal muscle	0.0	94743_Donor 3 U - B_Mesenchymal Stem Cells	0.0
97487_Patient-09ut_uterus	0.0	94730_Donor 3 AM - A_adipose	0.0
97488_Patient-09pl_placenta	0.0	94731_Donor 3 AM - B_adipose	0.0
97492_Patient-10ut_uterus	0.0	94732_Donor 3 AM - C_adipose	0.0
97493_Patient-10pl_placenta	0.0	94733_Donor 3 AD - A_adipose	0.0
97495_Patient-11go_adipose	0.0	94734_Donor 3 AD - B_adipose	0.0

97496_Patient-11sk_skeletal muscle	0.0	94735_Donor 3 AD - C_adipose	0.0
97497_Patient-11ut_uterus	0.0	77138_Liver_HepG2untreated	100.0
97498_Patient-11pl_placenta	0.0	73556_Heart_Cardiac stromal cells (primary)	0.0
97500_Patient-12go_adipose	0.0	81735_Small Intestine	25.5
97501_Patient-12sk_skeletal muscle	0.0	72409_Kidney_Proximal Convoluted Tubule	0.0
97502_Patient-12ut_uterus	0.0	82685_Small intestine_Duodenum	31.2
97503_Patient-12pl_placenta	0.0	90650_Adrenal_Adrenocortical adenoma	0.0
94721_Donor 2 U - A_Mesenchymal Stem Cells	0.0	72410_Kidney_HRCE	0.0
94722_Donor 2 U - B_Mesenchymal Stem Cells	0.0	72411_Kidney_HRE	0.0
94723_Donor 2 U - C_Mesenchymal Stem Cells	0.0	73139_Uterus_Uterine smooth muscle cells	0.0

CNS_neurodegeneration_v1.0 Summary: Ag6034 Expression of this gene is low/undetectable in all samples on this panel (CTs>35). (Data not shown.)

General_screening_panel_v1.5 Summary: Ag6034 Highest expression of this gene is seen in colon cancer (CT=26.3). High to moderate levels of expression are also seen in colon, renal, liver and lung cancer cell lines, as well as in fetal lung. This expression suggests that this gene may be involved in these cancers. Thus, expression of this gene could be used to differentiate between these samples and other samples on this panel and as a marker of these cancers. Therapeutic modulation of the expression or function of this gene may also be useful in the treatment of these cancers.

Panel 4.1D Summary: Ag6034 Expression of this gene is highest in colon and kidney (CTs=30). Thus, expression of this gene could be used as a marker of these tissues.

Panel 5 Islet Summary: Ag6034 Highest expression of this gene is seen in a liver cell line (CT=30.6). Thus, expression of this gene could be used to differentiate between this sample and other samples on this panel.

T. CG146513-01: Diacylglycerol acyltransferase 2.

Expression of gene CG146513-01 was assessed using the primer-probe set Ag6036, described in Table TA. Results of the RTQ-PCR runs are shown in Table TB.

Table TA. Probe Name Ag6036

Primers	Sequence	Length	Start Position	SEQ ID No
Forward	5'-tggaccctatggaagtatttcc-3'	22	326	324
Probe	TET-5'-ttcccagtacagctggtgaagactca-3'-TAMRA	26	356	325
Reverse	5'-gttgtgtttgggagaaagatca-3'	22	382	326

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Table TB. Panel 5 Islet

Tissue Name	Rel. Exp.(%) Ag603, Run 279370869	Tissue Name	Rel. Exp.(%) Ag6036, Run 279370869
97457_Patient-02go_adipose	10.5	94709_Donor 2 AM - A_adipose	11.4
97476_Patient-07sk_skeletal muscle	0.0	94710_Donor 2 AM - B_adipose	6.7
97477_Patient-07ut_uterus	3.3	94711_Donor 2 AM - C_adipose	4.2
97478_Patient-07pl_placenta	6.0	94712_Donor 2 AD - A_adipose	23.8
99167_Bayer Patient 1	3.3	94713_Donor 2 AD - B_adipose	32.8
97482_Patient-08ut_uterus	2.6	94714_Donor 2 AD - C_adipose	22.2
97483_Patient-08pl_placenta	1.0	94742_Donor 3 U - A_Mesenchymal Stem Cells	2.6
97486_Patient-09sk_skeletal muscle	8.4	94743_Donor 3 U - B_Mesenchymal Stem Cells	2.5
97487_Patient-09ut_uterus	5.8	94730_Donor 3 AM - A_adipose	12.9
97488_Patient-09pl_placenta	2.2	94731_Donor 3 AM - B_adipose	21.0
97492_Patient-10ut_uterus	4.0	94732_Donor 3 AM - C_adipose	20.4
97493_Patient-10pl_placenta	3.2	94733_Donor 3 AD - A_adipose	26.4
97495_Patient-11go_adipose	6.0	94734_Donor 3 AD - B_adipose	25.5
97496_Patient-11sk_skeletal muscle	20.2	94735_Donor 3 AD - C_adipose	6.5
97497_Patient-11ut_uterus	8.7	77138_Liver_HepG2untreated	41.5
97498_Patient-11pl_placenta	1.9	73556_Heart_Cardiac stromal cells (primary)	1.6
97500_Patient-12go_adipose	4.0	81735_Small Intestine	10.7
97501_Patient-12sk_skeletal muscle	22.2	72409_Kidney_Proximal Convoluted Tubule	100.0
97502_Patient-12ut_uterus	7.1	82685_Small intestine_Duodenum	15.7
97503_Patient-12pl_placenta	1.3	90650_Adrenal_Adrenocortical adenoma	5.0

94721_Donor 2 U - A_Mesenchymal Stem Cells	12.8	72410_Kidney_HRCE	31.2
94722_Donor 2 U - B_Mesenchymal Stem Cells	6.8	72411_Kidney_HRE	9.1
94723_Donor 2 U - C_Mesenchymal Stem Cells	11.2	73139_Uterus_Uterine smooth muscle cells	13.3

CNS_neurodegeneration_v1.0 Summary: Ag6036 Expression of this gene is low/undetectable in all samples on this panel (CTs>35).

General_screening_panel_v1.5 Summary: Ag6036 Expression of this gene is low/undetectable in all samples on this panel (CTs>35).

Panel 4.1D Summary: Ag6036 Expression of this gene is low/undetectable in all samples on this panel (CTs>35).

Panel 5 Islet Summary: Ag6036 Highest expression of this gene is seen in a kidney derived sample (CT=29.5). Moderate levels of expression are seen in many samples on this panel, including samples from uterus, placenta, adipose, and skeletal muscle. Thus, this gene may be involved in diseases of these tissues, including obesity and diabetes.

U. CG146522-01: Diacylglycerol acyltransferase 2.

Expression of gene CG146522-01 was assessed using the primer-probe set Ag6037, described in Table UA. Results of the RTQ-PCR runs are shown in Table UB.

Table UA. Probe Name Ag6037

Primers	Sequence	Length	Start Position	SEQ ID No
Forward	5'-attccaagcagcctagtcactt-3'	22	49	327
Probe	TET-5'-ttctgcagtggcctttgagctacctt-3'-TAMRA	26	85	328
Reverse	5'-cagcaggtagacgaacaagatg-3'	22	113	329

Table UB. Panel 5 Islet

Tissue Name	Rel. Exp.() Ag6037, Run 279370870	Tissue Name	Rel. Exp.(%) Ag6037, Run 279370870
97457_Patient-02go_adipose	0.0	94709_Donor 2 AM - A_adipose	0.0
97476_Patient-07sk_skeletal muscle	0.0	94710_Donor 2 AM - B_adipose	0.0
97477_Patient-07ut_uterus	0.0	94711_Donor 2 AM - C_adipose	0.0
97478_Patient-07pl_placenta	0.0	94712_Donor 2 AD - A_adipose	0.0
99167_Bayer Patient 1	0.9	94713_Donor 2 AD - B_adipose	0.0
97482_Patient-08ut_uterus	0.8	94714_Donor 2 AD - C_adipose	0.0
97483_Patient-08pl_placenta	0.0	94742_Donor 3 U - A_Mesenchymal Stem Cells	0.0
97486_Patient-09sk_skeletal muscle	9.0	94743_Donor 3 U - B_Mesenchymal Stem Cells	0.0
97487_Patient-09ut_uterus	2.2	94730_Donor 3 AM - A_adipose	0.0
97488_Patient-09pl_placenta	0.0	94731_Donor 3 AM - B_adipose	0.0
97492_Patient-10ut_uterus	0.5	94732_Donor 3 AM - C_adipose	0.0
97493_Patient-10pl_placenta	3.5	94733_Donor 3 AD - A_adipose	0.0
97495_Patient-11go_adipose	1.2	94734_Donor 3 AD - B_adipose	0.9
97496_Patient-11sk_skeletal muscle	39.2	94735_Donor 3 AD - C_adipose	0.0
97497_Patient-11ut_uterus	0.0	77138_Liver_HepG2untreated	0.0
97498_Patient-11pl_placenta	0.0	73556_Heart_Cardiac stromal cells (primary)	0.0
97500_Patient-12go_adipose	1.7	81735_Small Intestine	1.0
97501_Patient-12sk_skeletal muscle	**100.0**	72409_Kidney_Proximal Convoluted Tubule	0.0
97502_Patient-12ut_uterus	0.0	82685_Small intestine_Duodenum	0.0
97503_Patient-12pl_placenta	1.0	90650_Adrenal_Adrenocortical adenoma	0.0
94721_Donor 2 U - A_Mesenchymal Stem Cells	0.0	72410_Kidney_HRCE	0.0
94722_Donor 2 U - B_Mesenchymal Stem Cells	0.0	72411_Kidney_HRE	0.0
94723_Donor 2 U - C_Mesenchymal Stem Cells	0.5	73139_Uterus_Uterine smooth muscle cells	0.0

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CNS_neurodegeneration_v1.0 Summary: Ag6037 Expression of this gene is low/undetectable in all samples on this panel (CTs>35).

General_screening_panel_v1.5 Summary: Ag6037 Expression of this gene is
low/undetectable in all samples on this panel (CTs>35).

Panel 4.1D Summary: Ag6037 Expression of this gene is low/undetectable in all
samples on this panel (CTs>35).

Panel 5 Islet Summary: Ag6037 Expression of this gene is limited to skeletal
muscle (CTs=30-31). Thus, expression of this gene could be used to differentiate these
samples from other samples on this panel and as a marker of this tissue. Furthermore,
therapeutic modulation of the expression or function of this gene may be useful in the
treatment of metabolic disorders, including obesity and diabetes.

V. CG146531-01: DIACYLGLYCEROL ACYLTRANSFERASE 2.

Expression of gene CG146531-01 was assessed using the primer-probe set Ag6038,
described in Table VA.

Table VA. Probe Name Ag6038

Primers		Length	Start Position	SEQ ID No
Forward	5'-aaggtgtcacaggaagagcat-3'	21	10	330
Probe	TET-5'-agccaggtcaccatggctttcttct-3'-TAMRA	25	49	331
Reverse	5'-gccctcctggagattcagt-3'	19	78	332

CNS_neurodegeneration_v1.0 Summary: Ag6038 Expression of this gene is
low/undetectable in all samples on this panel (CTs>35).

General_screening_panel_v1.5 Summary: Ag6038 Expression of this gene is
low/undetectable in all samples on this panel (CTs>35).

Panel 4.1D Summary: Ag6038 Expression of this gene is low/undetectable in all
samples on this panel (CTs>35).

Panel 5 Islet Summary: Ag6038 Expression of this gene is low/undetectable in all
samples on this panel (CTs>35).

W. CG147274-01: Protease.

Expression of gene CG147274-01 was assessed using the primer-probe set Ag5623, described in Table WA.

Table WA. Probe Name Ag5623

Primers		Length	Start Position	SEQ ID No
Forward	5'-gatgtgctgccttcagaatg-3'	20	64	333
Probe	TET-5'-aatcctcccggcctccttggagt-3'-TAMRA	23	89	334
Reverse	5'-gtccttcctgggtgtcttg-3'	19	121	335

CNS_neurodegeneration_v1.0 Summary: Ag5623 Expression of this gene is low/undetectable in all samples on this panel (CTs>35).

General_screening_panel_v1.5 Summary: Ag5623 Expression of this gene is low/undetectable in all samples on this panel (CTs>35).

Panel 4.1D Summary: Ag5623 Expression of this gene is low/undetectable in all samples on this panel (CTs>35).

X. CG147419-01: GLUTAMINE: FRUCTOSE-6-PHOSPHATE AMIDOTRANSFERASE 1 MUSCLE.

Expression of gene CG147419-01 was assessed using the primer-probe set Ag5207, described in Table XA. Results of the RTQ-PCR runs are shown in Tables XB, XC, XD and XE.

Table XA. Probe Name Ag5207

Primers	Sequenes	Length	Start Position	SEQ ID No
Forward	5'-gccctctgttgattggtgta-3'	20	736	336
Probe	TET-5'-cggagtgaacataaactttctactgatca-3'-TAMRA	29	756	337
Reverse	5'-ccaatctgagtcctagctgttc-3'	22	802	338

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Table XB. CNS_neurodegeneration_v1.0

Tissue Name	Rel. Exp.(%) Ag5207, Run 226559656	issue Name	Rel. Exp.(%) Ag5207, Run 226559656
AD 1 Hippo	11.3	Control (Path) 3 Temporal Ctx	2.3
AD 2 Hippo	14.6	Control (Path) 4 Temporal Ctx	54.7
AD 3 Hippo	0.0	AD 1 Occipital Ctx	1.8
AD 4 Hippo	6.3	AD 2 Occipital Ctx (Missing)	0.0
AD 5 hippo	100.0	AD 3 Occipital Ctx	1.7
AD 6 Hippo	29.3	AD 4 Occipital Ctx	11.5
Control 2 Hippo	59.0	AD 5 Occipital Ctx	21.0
Control 4 Hippo	0.0	AD 6 Occipital Ctx	97.9
Control (Path) 3 Hippo	1.8	Control 1 Occipital Ctx	0.0
AD 1 Temporal Ctx	12.5	Control 2 Occipital Ctx	100.0
AD 2 Temporal Ctx	41.5	Control 3 Occipital Ctx	13.3
AD 3 Temporal Ctx	2.2	Control 4 Occipital Ctx	2.2
AD 4 Temporal Ctx	24.1	Control (Path) 1 Occipital Ctx	100.0
AD 5 Inf Temporal Ctx	65.5	Control (Path) 2 Occipital Ctx	7.2
AD 5 SupTemporal Ctx	29.1	Control (Path) 3 Occipital Ctx	0.0
AD 6 Inf Temporal Ctx	26.2	Control (Path) 4 Occipital Ctx	18.9
AD 6 Sup Temporal Ctx	49.3	Control 1 Parietal Ctx	2.5
Control 1 Temporal Ctx	0.0	Control 2 Parietal Ctx	53.2
Control 2 Temporal Ctx	88.3	Control 3 Parietal Ctx	21.6
Control 3 Temporal Ctx	19.5	Control (Path) 1 Parietal Ctx	94.6
Control 4 Temporal Ctx	4.9	Control (Path) 2 Parietal Ctx	16.8
Control (Path) 1 Temporal Ctx	97.3	Control (Path) 3 Parietal Ctx	4.0
Control (Path) 2 Temporal Ctx	48.0	Control (Path) 4 Parietal Ctx	50.3

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Table XC. General_screening_panel_v1.5

Tissue Name	Rel. Exp.(%) Ag5207, Run 228757767	issue Name	Rel. Exp.(%) Ag5207, Run 228757767
Adipose	9.9	Renal ca. TK-10	2.9
Melanoma* Hs688(A).T	4.0	Bladder	2.2
Melanoma* Hs688(B).T	12.1	Gastric ca. (liver met.) NCI-N87	23.2
Melanoma* M14	4.1	Gastric ca. KATO III	17.4
Melanoma* LOXIMVI	0.7	Colon ca. SW-948	0.4

Melanoma* SK-MEL-5	1.8	Colon ca. SW480	1.3
Squamous cell carcinoma SCC-4	0.7	Colon ca.* (SW480 met) SW620	0.1
Testis Pool	2.8	Colon ca. HT29	0.3
Prostate ca.* (bone met) PC-3	6.3	Colon ca. HCT-116	0.2
Prostate Pool	4.1	Colon ca. CaCo-2	1.6
Placenta	0.2	Colon cancer tissue	1.3
Uterus Pool	5.6	Colon ca. SW1116	0.0
Ovarian ca. OVCAR-3	0.2	Colon ca. Colo-205	2.6
Ovarian ca. SK-OV-3	5.9	Colon ca. SW-48	0.8
Ovarian ca. OVCAR-4	1.2	Colon Pool	11.3
Ovarian ca. OVCAR-5	1.6	Small Intestine Pool	4.2
Ovarian ca. IGROV-1	0.8	Stomach Pool	2.9
Ovarian ca. OVCAR-8	1.7	Bone Marrow Pool	2.1
Ovary	0.7	Fetal Heart	45.7
Breast ca. MCF-7	0.3	Heart Pool	38.2
Breast ca. MDA-MB-231	3.8	Lymph Node Pool	11.3
Breast ca. BT 549	1.3	Fetal Skeletal Muscle	19.3
Breast ca. T47D	0.0	Skeletal Muscle Pool	100.0
Breast ca. MDA-N	0.2	Spleen Pool	0.5
Breast Pool	6.4	Thymus Pool	4.0
Trachea	1.0	CNS cancer (glio/astro) U87-MG	11.0
Lung	1.5	CNS cancer (glio/astro) U-118-MG	24.0
Fetal Lung	1.2	CNS cancer (neuro;met) SK-N-AS	3.4
Lung ca. NCI-N417	0.7	CNS cancer (astro) SF-539	1.0
Lung ca. LX-1	0.6	CNS cancer (astro) SNB-75	1.4
Lung ca. NCI-H146	0.5	CNS cancer (glio) SNB-19	1.2
Lung ca. SHP-77	0.4	CNS cancer (glio) SF-295	18.6
Lung ca. A549	4.8	Brain (Amygdala) Pool	3.7
Lung ca. NCI-H526	0.6	Brain (cerebellum)	4.6
Lung ca. NCI-H23	0.2	Brain (fetal)	0.2
Lung ca. NCI-H460	3.2	Brain (Hippocampus) Pool	3.1
Lung ca. HOP-62	4.3	Cerebral Cortex Pool	6.7
Lung ca. NCI-H522	2.0	Brain (Substantia nigra) Pool	4.3
Liver	0.1	Brain (Thalamus) Pool	8.2
Fetal Liver	0.4	Brain (whole)	4.4
Liver ca. HepG2	0.4	Spinal Cord Pool	1.2
Kidney Pool	14.4	Adrenal Gland	2.6
Fetal Kidney	0.2	Pituitary gland Pool	1.5
Renal ca. 786-0	1.2	Salivary Gland	0.4
Renal ca. A498	1.2	Thyroid (female)	0.4
Renal ca. ACHN	1.6	Pancreatic ca. CAPAN2	2.8
Renal ca. UO-31	1.5	Pancreas Pool	6.0

Table XD. Panel 4.1D

Tissue Name	Rel. Ep.(%) Ag5207, Run 229739304	Tissue Name	Rel. Exp.(%) Ag5207, Run 229739304
Secondary Th1 act	0.0	HUVEC IL-1beta	16.0
Secondary Th2 act	4.2	HUVEC IFN gamma	9.6
Secondary Tr1 act	0.0	HUVEC TNF alpha + IFN gamma	3.5
Secondary Th1 rest	0.0	HUVEC TNF alpha + IL4	0.0
Secondary Th2 rest	0.0	HUVEC IL-11	5.5
Secondary Tr1 rest	0.0	Lung Microvascular EC none	7.1
Primary Th1 act	0.0	Lung Microvascular EC TNFalpha + IL-1beta	0.0
Primary Th2 act	5.6	Microvascular Dermal EC none	0.0
Primary Tr1 act	5.6	Microsvasular Dermal EC TNFalpha + IL-1beta	0.0
Primary Th1 rest	0.0	Bronchial epithelium TNFalpha + IL1beta	0.0
Primary Th2 rest	0.0	Small airway epithelium none	5.8
Primary Tr1 rest	0.0	Small airway epithelium TNFalpha + IL-1beta	3.6
CD45RA CD4 lymphocyte act	35.6	Coronery artery SMC rest	7.4
CD45RO CD4 lymphocyte act	0.0	Coronery artery SMC TNFalpha + IL-1beta	13.6
CD8 lymphocyte act	7.8	Astrocytes rest	0.0
Secondary CD8 lymphocyte rest	0.0	Astrocytes TNFalpha + IL-1beta	12.9
Secondary CD8 lymphocyte act	0.0	KU-812 (Basophil) rest	0.0
CD4 lymphocyte none	10.7	KU-812 (Basophil) PMA/ionomycin	0.0
2ry Th1/Th2/Tr1_anti-CD95 CH11	0.0	CCD1106 (Keratinocytes) none	18.6
LAK cells rest	0.0	CCD1106 (Keratinocytes) TNFalpha + IL-1beta	0.0
LAK cells IL-2	0.0	Liver cirrhosis	0.0
LAK cells IL-2+IL-12	0.0	NCI-H292 none	0.0
LAK cells IL-2+IFN gamma	0.0	NCI-H292 IL-4	0.0
LAK cells IL-2+ IL-18	0.0	NCI-H292 IL-9	0.0
LAK cells PMA/ionomycin	6.0	NCI-H292 IL-13	0.0
NK Cells IL-2 rest	4.8	NCI-H292 IFN gamma	0.0
Two Way MLR 3 day	0.0	HPAEC none	4.7
Two Way MLR 5 day	0.0	HPAEC TNF alpha + IL-1 beta	20.9
Two Way MLR 7 day	0.0	Lung fibroblast none	17.7
PBMC rest	7.0	Lung fibroblast TNF alpha + IL-1 beta	23.0

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PBMC PWM	0.0	Lung fibroblast IL-4	10.8
PBMC PHA-L	0.0	Lung fibroblast IL-9	11.3
Ramos (B cell) none	0.0	Lung fibroblast IL-13	9.2
Ramos (B cell) ionomycin	0.0	Lung fibroblast IFN gamma	33.0
B lymphocytes PWM	0.0	Dermal fibroblast CCD1070 rest	41.8
B lymphocytes CD40L and IL-4	0.0	Dermal fibroblast CCD1070 TNF alpha	100.0
EOL-1 dbcAMP	0.0	Dermal fibroblast CCD1070 IL-1 beta	77.9
EOL-1 dbcAMP PMA/ionomycin	0.0	Dermal fibroblast IFN gamma	7.6
Dendritic cells none	5.1	Dermal fibroblast IL-4	15.3
Dendritic cells LPS	0.0	Dermal Fibroblasts rest	34.6
Dendritic cells anti-CD40	0.0	Neutrophils TNFa+LPS	4.8
Monocytes rest	6.6	Neutrophils rest	0.0
Monocytes LPS	0.0	Colon	0.0
Macrophages rest	0.0	Lung	12.3
Macrophages LPS	0.0	Thymus	0.0
HUVEC none	6.0	Kidney	0.0
HUVEC starved	29.5		

Table XE. Panel 5 Islet

5

Tissue Name	Rel. Exp.() Ag5207, Run 263594763	Tissue Name	Rel. Exp.(%) Ag5207, Run 263594763
97457_Patient-02go_adipose	2.0	94709_Donor 2 AM - A_adipose	4.6
97476_Patient-07sk_skeletal muscle	3.1	94710_Donor 2 AM - B_adipose	1.1
97477_Patient-07ut_uterus	3.2	94711_Donor 2 AM - C_adipose	0.8
97478_Patient-07pl_placenta	2.0	94712_Donor 2 AD - A_adipose	1.0
99167_Bayer Patient 1	1.0	94713_Donor 2 AD - B_adipose	8.1
97482_Patient-08ut_uterus	6.7	94714_Donor 2 AD - C_adipose	5.3
97483_Patient-08pl_placenta	0.0	94742_Donor 3 U - A_Mesenchymal Stem Cells	1.2
97486_Patient-09sk_skeletal muscle	27.4	94743_Donor 3 U - B_Mesenchymal Stem Cells	3.7
97487_Patient-09ut_uterus	12.4	94730_Donor 3 AM - A_adipose	4.6
97488_Patient-09pl_placenta	1.3	94731_Donor 3 AM - B_adipose	2.1
97492_Patient-10ut_uterus	14.4	94732_Donor 3 AM - C_adipose	1.0
97493_Patient-10pl_placenta	2.1	94733_Donor 3 AD - A_adipose	6.9
97495_Patient-11go_adipose	2.0	94734_Donor 3 AD - B_adipose	3.2

97496_Patient-11sk_skeletal muscle	50.3	94735_Donor 3 AD - C_adipose	4.4
97497_Patient-11ut_uterus	7.1	77138_Liver_HepG2untreated	3.4
97498_Patient-11pl_placenta	0.0	73556_Heart_Cardiac stromal cells (primary)	2.2
97500_Patient-12go_adipose	10.7	81735_Small Intestine	7.1
97501_Patient-12sk_skeletal muscle	100.0	72409_Kidney_Proximal Convoluted Tubule	0.0
97502_Patient-12ut_uterus	10.9	82685_Small intestine_Duodenum	0.0
97503_Patient-12pl_placenta	0.0	90650_Adrenal_Adrenocortical adenoma	0.0
94721_Donor 2 U - A_Mesenchymal Stem Cells	1.8	72410_Kidney_HRCE	4.9
94722_Donor 2 U - B_Mesenchymal Stem Cells	1.0	72411_Kidney_HRE	0.0
94723_Donor 2 U - C_Mesenchymal Stem Cells	3.5	73139_Uterus_Uterine smooth muscle cells	4.0

CNS_neurodegeneration_v1.0 Summary: Ag5207 This panel does not show differential expression of this gene in Alzheimer's disease. However, this profile confirms the expression of this gene at moderate levels in the brain. Please see Panel 1.5 for discussion of this gene in the central nervous system.

General_screening_panel_v1.5 Summary: Ag5207 Highest expression of this gene is seen in skeletal muscle (CT=28). Low but significant expression is also seen in pancreas, adrenal, pituitary, adipose, adult and fetal heart, and fetal skeletal muscle. This gene encodes a protein that is homologous to Glutamine:fructose-6-phosphate amidotransferase (GFAT) which catalyzes the formation of glucosamine 6-phosphate and is the first and rate-limiting enzyme of the hexosamine biosynthetic pathway. Enhanced glucose flux via the hexosamine biosynthetic pathway has been implicated in in the induction of insulin resistance. Buse et al. showed in a mouse model that glucose flux via the hexosamine pathway is selectively increased in muscle and may contribute to muscle insulin resistance in non-insulin-dependent diabetes mellitus. (Am J Physiol 1997 Jun;272(6 Pt 1):E1080-8). Thus, based on the homology of this enzyme to GFAT and the high expression in muscle, modulation of the expression or function of this gene may be useful in the treatment of type II diabetes.

This gene is widely expressed on this panel with moderate to low expression seen throughout the CNS, including the hippocampus, thalamus, substantia nigra, amygdala, cerebellum and cerebral cortex. Therefore, therapeutic modulation of the expression or

function of this gene may be useful in the treatment of neurological disorders, such as Alzheimer's disease, Parkinson's disease, schizophrenia, multiple sclerosis, stroke and epilepsy.

Moderate to low levels of expression are also seen in many cancer cell lines on this panel, including gastric cancer and melanoma cell lines. Thus, modulation of this gene product may be useful in the treatment of cancer.

Panel 4.1D Summary: Ag5207 Detectable levels of expression appear to be restricted to TNF-alpha treated dermal fibroblasts (CT=33.3). This expression suggests that this gene product may be involved in skin disorders, including psoriasis.

Panel 5 Islet Summary: Ag5207 Highest expression is seen in skeletal muscle (CT=30.2), in agreement with panel 1.5. Moderate to low levels of expression are also seen in other metabolic tissues, including uterus and adipose. Please see Panel 1.5 for discussion of this gene in metabolic disease.

Y. CG148102-01: CARNITINE O-PALMITOYLTRANSFERASE I.

Expression of gene CG148102-01 was assessed using the primer-probe set Ag5274, described in Table YA. Results of the RTQ-PCR runs are shown in Tables YB, YC, YD and YE.

Table YA. Probe Name Ag5274

Primers		Length	Start Position	SEQ ID No
Forward	5'-cacttccgggacccacagt-3'	19	1732	339
Probe	TET-5'-caccaggctctgctgaaggcagcc-3'-TAMRA	24	1783	340
Reverse	5'-caaacaggtggcggtcaact-3'	20	1821	341

Table YB. CNS_neurodegeneration_v1.0

Tissue Name	Rel. Exp.(%) Ag5274, Run 230512893	Tissue Name	Rel. Exp.(%) Ag5274, Run 230512893

AD 1 Hippo	19.3	Control (Path) 3 Temporal Ctx	7.7
AD 2 Hippo	33.2	Control (Path) 4 Temporal Ctx	29.7
AD 3 Hippo	11.7	AD 1 Occipital Ctx	18.3
AD 4 Hippo	9.9	AD 2 Occipital Ctx (Missing)	0.0
AD 5 Hippo	95.9	AD 3 Occipital Ctx	7.5
AD 6 Hippo	43.5	AD 4 Occipital Ctx	15.1
Control 2 Hippo	57.0	AD 5 Occipital Ctx	66.4
Control 4 Hippo	11.9	AD 6 Occipital Ctx	13.1
Control (Path) 3 Hippo	8.5	Control 1 Occipital Ctx	3.7
AD 1 Temporal Ctx	17.0	Control 2 Occipital Ctx	98.6
AD 2 Temporal Ctx	29.5	Control 3 Occipital Ctx	27.5
AD 3 Temporal Ctx	8.3	Control 4 Occipital Ctx	4.5
AD 4 Temporal Ctx	19.6	Control (Path) 1 Occipital Ctx	**100.0**
AD 5 Inf Temporal Ctx	95.9	Control (Path) 2 Occipital Ctx	17.1
AD 5 Sup Temporal Ctx	53.6	Control (Path) 3 Occipital Ctx	3.8
AD 6 Inf Temporal Ctx	29.9	Control (Path) 4 Occipital Ctx	20.0
AD 6 Sup Temporal Ctx	33.2	Control 1 Parietal Ctx	10.5
Control 1 Temporal Ctx	8.4	Control 2 Parietal Ctx	49.3
Control 2 Temporal Ctx	70.2	Control 3 Parietal Ctx	19.2
Control 3 Temporal Ctx	25.0	Control (Path) 1 Parietal Ctx	94.6
Control 3 Temporal Ctx	11.3	Control (Path) 2 Parietal Ctx	25.0
Control (Path) 1 Temporal Ctx	74.2	Control (Path) 3 Parietal Ctx	6.0
Control (Path) 2 Temporal Ctx	44.4	Control (Path) 4 Parietal Ctx	50.7

Table YC. General_screening_panel_v1.5

5

Tissue Name	Rel. Exp.(%) Ag5274, Run 230762793	Tissue Name	Rel. Exp.(%) Ag5274, Run 230762793
Adipose	1.2	Renal ca. TK-10	0.0
Melanoma* Hs688(A).T	7.4	Bladder	1.7
Melanoma* Hs688(B).T	13.0	Gastric ca. (liver met.) NCI-N87	1.0
Melanoma* M14	0.1	Gastric ca. KATO III	0.2
Melanoma* LOXIMVI	0.0	Colon ca. SW-948	1.4
Melanoma* SK-MEL-5	0.0	Colon ca. SW480	0.7
Squamous cell carcinoma SCC-4	1.5	Colon ca.* (SW480 met) SW620	0.0
Testis Pool	2.1	Colon ca. HT29	0.2
Prostate ca.* (bone met) PC-3	21.8	Colon ca. HCT-116	2.1
Prostate Pool	0.8	Colon ca. CaCo-2	0.3
Placenta	0.7	Colon cancer tissue	2.4
Uterus Pool	0.7	Colon ca. SW1116	0.0

Ovarian ca. OVCAR-3	12.2	Colon ca. Colo-205	0.0
Ovarian ca. SK-OV-3	0.2	Colon ca. SW-48	0.0
Ovarian ca. OVCAR-4	0.1	Colon Pool	3.5
Ovarian ca. OVCAR-5	2.8	Small Intestine Pool	2.1
Ovarian ca. IGROV-1	7.2	Stomach Pool	1.8
Ovarian ca. OVCAR-8	3.9	Bone Marrow Pool	0.8
Ovary	6.3	Fetal Heart	1.7
Breast ca. MCF-7	0.2	Heart Pool	1.5
Breast ca. MDA-MB-231	4.9	Lymph Node Pool	5.3
Breast ca. BT 549	88.3	Fetal Skeletal Muscle	1.0
Breast ca. T47D	0.0	Skeletal Muscle Pool	0.8
Breast ca. MDA-N	0.0	Spleen Pool	3.0
Breast Pool	4.9	Thymus Pool	2.7
Trachea	1.0	CNS cancer (glio/astro) U87-MG	27.7
Lung	0.9	CNS cancer (glio/astro) U-118-MG	27.4
Fetal Lung	7.2	CNS cancer (neuro;met) SK-N-AS	86.5
Lung ca. NCI-N417	8.2	CNS cancer (astro) SF-539	0.0
Lung ca. LX-1	0.5	CNS cancer (astro) SNB-75	0.5
Lung ca. NCI-H146	16.2	CNS cancer (glio) SNB-19	7.2
Lung ca. SHP-77	53.6	CNS cancer (glio) SF-295	17.3
Lung ca. A549	0.0	Brain (Amygdala) Pool	19.9
Lung ca. NCI-H526	3.6	Brain (cerebellum)	100.0
Lung ca. NCI-H23	40.9	Brain (fetal)	44.8
Lung ca. NCI-H460	0.6	Brain (Hippocampus) Pool	16.8
Lung ca. HOP-62	1.6	Cerebral Cortex Pool	24.0
Lung ca. NCI-H522	57.8	Brain (Substantia nigra) Pool	27.4
Liver	0.3	Brain (Thalamus) Pool	34.2
Fetal Liver	0.9	Brain (whole)	42.0
Liver ca. HepG2	0.0	Spinal Cord Pool	10.5
Kidney Pool	4.2	Adrenal Gland	1.0
Fetal Kidney	3.6	Pituitary gland Pool	4.9
Renal ca. 786-0	0.0	Salivary Gland	0.1
Renal ca. A498	0.0	Thyroid (female)	0.6
Renal ca. ACHN	0.5	Pancreatic ca. CAPAN2	0.0
Renal ca. UO-31	0.3	Pancreas Pool	4.8

Table YD. Panel 4.1D

5

Tissue Name	Rel. Ep.(%) Ag5274, Run 230472159	Tissue Name	Rel. Exp.(%) Ag5274, Run 230472159

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Secondary Th1 act	2.3	HUVEC IL-1beta	45.1
Secondary Th2 act	1.6	HUVEC IFN gamma	92.0
Secondary Tr1 act	0.0	HUVEC TNF alpha + IFN gamma	15.1
Secondary Th1 rest	0.0	HUVEC TNF alpha + IL4	11.7
Secondary Th2 rest	2.3	HUVEC IL-11	67.8
Secondary Tr1 rest	0.0	Lung Microvascular EC none	38.2
Primary Th1 act	0.0	Lung Microvascular EC TNFalpha + IL-1beta	9.2
Primary Th2 act	0.0	Microvascular Dermal EC none	26.2
Primary Tr1 act	0.0	Microsvasular Dermal EC TNFalpha + IL-1beta	9.0
Primary Th1 rest	0.0	Bronchial epithelium TNFalpha + IL1beta	0.0
Primary Th2 rest	4.6	Small airway epithelium none	0.0
Primary Tr1 rest	0.0	Small airway epithelium TNFalpha + IL-1beta	0.0
CD45RA CD4 lymphocyte act	7.8	Coronery artery SMC rest	56.6
CD45RO CD4 lymphocyte act	0.0	Coronery artery SMC TNFalpha + IL-1beta	66.9
CD8 lymphocyte act	0.0	Astrocytes rest	23.2
Secondary CD8 lymphocyte rest	0.0	Astrocytes TNFalpha + IL-1beta	14.8
Secondary CD8 lymphocyte act	0.0	KU-812 (Basophil) rest	0.0
CD4 lymphocyte none	0.0	KU-812 (Basophil) PMA/ionomycin	0.0
2ry Th1/Th2/Tr1_anti-CD95 CH11	0.0	CCD1106 (Keratinocytes) none	31.9
LAK cells rest	0.0	CCD1106 (Keratinocytes) TNFalpha + IL-1beta	9.4
LAK cells IL-2	0.0	Liver cirrhosis	5.1
LAK cells IL-2+IL-12	0.0	NCI-H292 none	0.0
LAK cells IL-2+IFN gamma	0.0	NCI-H292 IL-4	0.0
LAK cells IL-2+ IL-18	0.0	NCI-H292 IL-9	0.0
LAK cells PMA/ionomycin	0.0	NCI-H292 IL-13	0.0
NK Cells IL-2 rest	2.5	NCI-H292 IFN gamma	8.6
Two Way MLR 3 day	0.0	HPAEC none	45.4
Two Way MLR 5 day	0.0	HPAEC TNF alpha + IL-1 beta	27.9
Two Way MLR 7 day	0.0	Lung fibroblast none	100.0
PBMC rest	0.0	Lung fibroblast TNF alpha + IL-1 beta	90.8
PBMC PWM	2.2	Lung fibroblast IL-4	22.2
PBMC PHA-L	10.1	Lung fibroblast IL-9	47.6
Ramos (B cell) none	0.0	Lung fibroblast IL-13	11.8
Ramos (B cell) ionomycin	0.0	Lung fibroblast IFN gamma	61.1
B lymphocytes PWM	0.0	Dermal fibroblast CCD1070 rest	28.7
B lymphocytes CD40L and IL-4	2.2	Dermal fibroblast CCD1070 TNF alpha	23.3

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EOL-1 dbcAMP	9.2	Dermal fibroblast CCD1070 IL-1 beta	28.7
EOL-1 dbcAMP PMA/ionomycin	2.7	Dermal fibroblast IFN gamma	16.7
Dendritic cells none	0.0	Dermal fibroblast IL-4	13.1
Dendritic cells LPS	0.0	Dermal Fibroblasts rest	58.6
Dendritic cells anti-CD40	0.0	Neutrophils TNFa+LPS	0.0
Monocytes rest	0.0	Neutrophils rest	0.0
Monocytes LPS	0.0	Colon	0.0
Macrophages rest	0.0	Lung	1.7
Macrophages LPS	0.0	Thymus	0.0
HUVEC none	48.3	Kidney	5.5
HUVEC starved	61.1		

Table YE. Panel 5 Islet

Tissue Name	Rel. Exp.() Ag5274, Run 307720339	Tissue Name	Rel. Exp.(%) Ag5274, Run 307720339
97457_Patient-02go_adipose	15.3	94709_Donor 2 AM - A_adipose	13.9
97476_Patient-07sk_skeletal muscle	0.0	94710_Donor 2 AM - B_adipose	15.2
97477_Patient-07ut_uterus	13.7	94711_Donor 2 AM - C_adipose	19.8
97478_Patient-07pl_placenta	9.0	94712_Donor 2 AD - A_adipose	58.2
99167_Bayer Patient 1	51.8	94713_Donor 2 AD - B_adipose	29.7
97482_Patient-08ut_uterus	24.3	94714_Donor 2 AD - C_adipose	34.9
97483_Patient-08pl_placenta	0.0	94742_Donor 3 U - A_Mesenchymal Stem Cells	62.9
97486_Patient-09sk_skeletal muscle	0.0	94743_Donor 3 U - B_Mesenchymal Stem Cells	39.5
97487_Patient-09ut_uterus	7.3	94730_Donor 3 AM - A_adipose	31.4
97488_Patient-09pl_placenta	11.9	94731_Donor 3 AM - B_adipose	35.1
97492_Patient-10ut_uterus	12.8	94732_Donor 3 AM - C_adipose	49.3
97493_Patient-10pl_placenta	5.3	94733_Donor 3 AD - A_adipose	28.9
97495_Patient-11go_adipose	5.3	94734_Donor 3 AD - B_adipose	44.8
97496_Patient-11sk_skeletal muscle	3.8	94735_Donor 3 AD - C_adipose	17.7
97497_Patient-11ut_uterus	20.9	77138_Liver_HepG2untreated	6.0
97498_Patient-11pl_placenta	5.4	73556_Heart_Cardiac stromal cells (primary)	55.5
97500_Patient-12go_adipose	27.0	81735_Small Intestine	39.0

97501_Patient-12sk_skeletal muscle	12.5	72409_Kidney_Proximal Convoluted Tubule	15.2
97502_Patient-12ut_uterus	10.2	82685_Small intestine_Duodenum	0.0
97503_Patient-12pl_placenta	2.4	90650_Adrenal_Adrenocortical adenoma	12.2
94721_Donor 2 U - A_Mesenchymal Stem Cells	100.0	72410_Kidney_HRCE	0.0
94722_Donor 2 U - B_Mesenchymal Stem Cells	43.2	72411_Kidney_HRE	25.7
94723_Donor 2 U - C_Mesenchymal Stem Cells	63.7	73139_Uterus_Uterine smooth muscle cells	97.9

CNS_neurodegeneration_v1.0 Summary: Ag5274 This panel confirms the expression of this gene at low levels in the brain in an independent group of individuals. This gene appears to be slightly down-regulated in the temporal cortex of Alzheimer's disease patients. Therefore, up-regulation of this gene or its protein product, or treatment with specific agonists for this receptor may be of use in reversing the dementia, memory loss, and neuronal death associated with this disease.

General_screening_panel_v1.5 Summary: Ag5274 Highest expression of this gene is seen in the cerebellum (CT=29.3). Moderate expression of this gene is seen throughout the brain. Thus, this gene would be useful for distinguishing brain tissue from non-neural tissue, and may be beneficial as a drug target in neurodegenerative disease, and specifically disorders that have this brain region as the site of pathology, such as autism and the ataxias. Please see Panel_CNS_neurodegeneration for further discussion of potential utility in the central nervous system.

Low but significant expression is also seen in pancreas. This gene encodes a protein with homology to carnitine palmitoyltransferase. Giannessi et al has shown that inhibition of this enzyme produces a significant reduction in serum glucose levels (J Med Chem 2001 Jul 19;44(15):2383-6). Thus, modulation of this enzyme may also be useful in the treatment of obesity and/or diabetes.

Panel 4.1D Summary: Ag5274 Highest expression of this gene is seen in untreated lung fibroblasts. Low, but significant expression is also seen in a cluster of treated and untreated lung and dermal fibroblasts. Low levels of expression are also seen in coronary artery SMCs, and HUVECs. This profile suggests that this gene could be used to differentiate between these cells and other cells samples. In addition, this gene product may be involved in inflammatory conditions of the lung and skin.

Panel 5 Islet Summary: Ag5274 Expression is limited to a sample derived from mesenchymal stem cells (CTs=34.5).

Z. CG148431-01 and CG148431-02: AMINOTRANSFERASE SIMILAR TO SERINE PALMOTYLTRANSFERASE.

5   Expression of gene CG148431-01 and CG148431-02 was assessed using the primer-probe set Ag5627, described in Table ZA. Results of the RTQ-PCR runs are shown in Tables ZB, ZC, ZD and ZE. Please note that CG148431-02 represents a full-length physical clone of the CG148431-01 gene, validating the prediction of the gene sequence.

Table ZA. Probe Name Ag5627
10

Primers	Sequenes	Length	Start Position	SEQ ID No
Forward	5'-gggctcctataacttccttggt-3'	22	555	342
Probe	TET-5'-tcctcatagactcatcatacttggctgca-3'-TAMRA	29	579	343
Reverse	5'-cctgtgccatacacctctaaaa-3'	22	620	344

Table ZB. CNS_neurodegeneration_v1.0
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Tissue Name	Rel. Exp.(%) Ag5627, Run 246956910	Rel. Exp.(%) Ag5627, Run 264979289	issue Name	Rel. Exp.(%) Ag5627, Run 246956910	Rel. Exp.(%) Ag5627, Run 264979289
AD 1 Hippo	17.4	57.0	Control (Path) 3 Temporal Ctx	6.4	8.2
AD 2 Hippo	67.8	4.8	Control (Path) 4 Temporal Ctx	10.3	24.0
AD 3 Hippo	50.0	62.4	AD 1 Occipital Ctx	11.8	26.8
AD 4 Hippo	19.1	30.8	AD 2 Occipital Ctx (Missing)	0.0	0.0
AD 5 Hippo	17.0	31.2	AD 3 Occipital Ctx	4.2	25.9
AD 6 Hippo	100.0	86.5	AD 4 Occipital Ctx	20.0	27.9
Control 2 Hippo	24.1	31.6	AD 5 Occipital Ctx	37.4	17.0
Control 4 Hippo	50.7	70.7	AD 6 Occipital Ctx	29.1	22.4
Control (Path) 3 Hippo	21.0	24.3	Control 1 Occipital Ctx	3.9	12.1
AD 1 Temporal Ctx	43.8	65.5	Control 2 Occipital Ctx	20.6	29.9

AD 2 Temporal Ctx	47.6	**100.0**	Control 3 Occipital Ctx	9.3	19.9
AD 3 Temporal Ctx	11.0	23.0	Control 4 Occipital Ctx	16.3	44.1
AD 4 Temporal Ctx	20.4	33.9	Control (Path) 1 Occipital Ctx	49.0	58.2
AD 5 Inf Temporal Ctx	31.0	31.2	Control (Path) 2 Occipital Ctx	6.6	15.2
AD 5 Sup Temporal Ctx	51.1	63.3	Control (Path) 3 Occipital Ctx	0.0	1.6
AD 6 Inf Temporal Ctx	68.8	87.7	Control (Path) 4 Occipital Ctx	23.3	14.3
AD 6 Sup Temporal Ctx	56.3	97.3	Control 1 Parietal Ctx	13.1	18.3
Control 1 Temporal Ctx	7.3	4.5	Control 2 Parietal Ctx	31.6	68.8
Control 2 Temporal Ctx	12.9	31.6	Control 3 Parietal Ctx	7.9	19.8
Control 3 Temporal Ctx	7.9	15.0	Control (Path) 1 Parietal Ctx	63.7	87.1
Control 3 Temporal Ctx	13.8	15.6	Control (Path) 2 Parietal Ctx	51.1	57.4
Control (Path) 1 Temporal Ctx	30.1	46.0	Control (Path) 3 Parietal Ctx	3.1	6.1
Control (Path) 2 Temporal Ctx	28.7	39.5	Control (Path) 4 Parietal Ctx	54.7	59.5

Table ZC. Panel 4.1D

Tissue Name	Rel. Ep.(%) Ag5627, Run 246490777	Tissue Name	Rel. Exp.(%) Ag5627, Run 246490777
Secondary Th1 act	0.0	HUVEC IL-1beta	0.0
Secondary Th2 act	0.4	HUVEC IFN gamma	16.7
Secondary Tr1 act	0.0	HUVEC TNF alpha + IFN gamma	0.3
Secondary Th1 rest	0.0	HUVEC TNF alpha + IL4	0.0
Secondary Th2 rest	0.0	HUVEC IL-11	1.2
Secondary Tr1 rest	0.0	Lung Microvascular EC none	0.4
Primary Th1 act	0.0	Lung Microvascular EC TNFalpha + IL-1beta	0.0
Primary Th2 act	0.2	Microvascular Dermal EC none	0.0
Primary Tr1 act	0.2	Microsvasular Dermal EC TNFalpha + IL-1beta	0.0

Primary Th1 rest	0.0	Bronchial epithelium TNFalpha + IL1beta	8.4
Primary Th2 rest	0.0	Small airway epithelium none	18.7
Primary Tr1 rest	0.0	Small airway epithelium TNFalpha + IL-1beta	24.3
CD45RA CD4 lymphocyte act	2.7	Coronery artery SMC rest	3.3
CD45RO CD4 lymphocyte act	6.8	Coronery artery SMC TNFalpha + IL-1beta	2.8
CD8 lymphocyte act	0.0	Astrocytes rest	3.9
Secondary CD8 lymphocyte rest	0.8	Astrocytes TNFalpha + IL-1beta	1.4
Secondary CD8 lymphocyte act	0.0	KU-812 (Basophil) rest	8.0
CD4 lymphocyte none	0.0	KU-812 (Basophil) PMA/ionomycin	14.2
2ry Th1/Th2/Tr1_anti-CD95 CH11	0.4	CCD1106 (Keratinocytes) none	17.4
LAK cells rest	0.0	CCD1106 (Keratinocytes) TNFalpha + IL-1beta	24.3
LAK cells IL-2	0.0	Liver cirrhosis	13.3
LAK cells IL-2+IL-12	0.2	NCI-H292 none	10.2
LAK cells IL-2+IFN gamma	0.0	NCI-H292 IL-4	36.3
LAK cells IL-2+ IL-18	0.0	NCI-H292 IL-9	21.5
LAK cells PMA/ionomycin	0.2	NCI-H292 IL-13	27.7
NK Cells IL-2 rest	11.8	NCI-H292 IFN gamma	18.3
Two Way MLR 3 day	0.4	HPAEC none	0.8
Two Way MLR 5 day	0.0	HPAEC TNF alpha + IL-1 beta	0.3
Two Way MLR 7 day	0.0	Lung fibroblast none	21.5
PBMC rest	0.0	Lung fibroblast TNF alpha + IL-1 beta	2.7
PBMC PWM	0.0	Lung fibroblast IL-4	10.2
PBMC PHA-L	1.3	Lung fibroblast IL-9	6.2
Ramos (B cell) none	0.0	Lung fibroblast IL-13	1.3
Ramos (B cell) ionomycin	0.0	Lung fibroblast IFN gamma	43.5
B lymphocytes PWM	0.0	Dermal fibroblast CCD1070 rest	0.0
B lymphocytes CD40L and IL-4	0.0	Dermal fibroblast CCD1070 TNF alpha	1.1
EOL-1 dbcAMP	3.5	Dermal fibroblast CCD1070 IL-1 beta	1.6
EOL-1 dbcAMP PMA/ionomycin	0.0	Dermal fibroblast IFN gamma	39.5
Dendritic cells none	1.1	Dermal fibroblast IL-4	12.0
Dendritic cells LPS	0.0	Dermal Fibroblasts rest	16.0
Dendritic cells anti-CD40	0.0	Neutrophils TNFa+LPS	0.0
Monocytes rest	0.0	Neutrophils rest	0.0
Monocytes LPS	0.0	Colon	3.0
Macrophages rest	0.0	Lung	4.6
Macrophages LPS	0.0	Thymus	3.5

HUVEC none	0.7	Kidney	100.0
HUVEC starved	2.9		

Table ZD. Panel 5 Islet

Tissue Name	Rel. Exp.(%) Ag5627 Run 279371483	Rel. Exp.(%) Ag5627, Run 312852505	Tissue Name	Rel. Exp.(%) Ag5627, Run 279371483	Rel. Exp.(%) Ag5627, Run 312852505
97457_Patient-02go_adipose	0.7	1.7	94709_Donor 2 AM - A_adipose	1.2	1.6
97476_Patient-07sk_skeletal muscle	0.0	0.0	94710_Donor 2 AM - B_adipose	1.1	1.7
97477_Patient-07ut_uterus	0.4	0.5	94711_Donor 2 AM - C_adipose	0.8	1.4
97478_Patient-07pl_placenta	40.3	46.0	94712_Donor 2 AD - A_adipose	2.7	2.0
99167_Bayer Patient 1	0.1	0.1	94713_Donor 2 AD - B_adipose	4.0	3.0
97482_Patient-08ut_uterus	0.2	0.2	94714_Donor 2 AD - C_adipose	3.0	3.0
97483_Patient-08pl_placenta	82.9	**100.0**	94742_Donor 3 U - A_Mesenchymal Stem Cells	0.4	0.4
97486_Patient-09sk_skeletal muscle	0.2	0.1	94743_Donor 3 U - B_Mesenchymal Stem Cells	0.3	0.6
97487_Patient-09ut_uterus	0.2	0.5	94730_Donor 3 AM - A_adipose	3.5	3.7
97488_Patient-09pl_placenta	29.9	25.5	94731_Donor 3 AM - B_adipose	5.3	5.6
97492_Patient-10ut_uterus	0.3	0.4	94732_Donor 3 AM - C_adipose	3.9	4.8
97493_Patient-10pl_placenta	100.0	71.7	94733_Donor 3 AD - A_adipose	2.6	3.5
97495_Patient-11go_adipose	1.2	0.9	94734_Donor 3 AD - B_adipose	2.8	3.6
97496_Patient-11sk_skeletal muscle	0.2	0.1	94735_Donor 3 AD - C_adipose	0.5	0.8
97497_Patient-11ut_uterus	0.5	0.8	77138_Liver_HepG2untreated	39.5	43.2
97498_Patient-11pl_placenta	28.1	31.6	73556_Heart_Cardiac stromal cells (primary)	0.1	0.0
97500_Patient-12go_adipose	1.0	1.8	81735_Small Intestine	1.8	1.9
97501_Patient-12sk_skeletal muscle	0.5	0.6	72409_Kidney_Proximal Convoluted Tubule	18.2	19.1

97502_Patient-12ut_uterus	0.3	0.4	82685_Small intestine_Duodenum	1.3	1.1
97503_Patient-12pl_placenta	85.9	88.3	90650_Adrenal_Adrenocortical adenoma	0.6	0.4
94721_Donor 2 U - A_Mesenchymal Stem Cells	1.2	1.3	72410_Kidney_HRCE	3.7	4.9
94722_Donor 2 U - B_Mesenchymal Stem Cells	0.6	0.8	72411_Kidney_HRE	1.6	1.7
94723_Donor 2 U - C_Mesenchymal Stem Cells	1.0	1.3	73139_Uterus_Uterine smooth muscle cells	1.0	0.7

Table ZE. general oncology screening panel_v_2.4

Tissue Name	Rel. Exp.(%) Ag5627, Run 268787222	Tissue ame	Rel. Exp.(%) Ag5627, Run 268787222
Colon cancer 1	2.8	Bladder NAT 2	0.3
Colon NAT 1	2.7	Bladder NAT 3	0.2
Colon cancer 2	7.8	Bladder NAT 4	1.1
Colon NAT 2	3.1	Prostate adenocarcinoma 1	11.8
Colon cancer 3	5.7	Prostate adenocarcinoma 2	1.0
Colon NAT 3	6.4	Prostate adenocarcinoma 3	8.6
Colon malignant cancer 4	3.0	Prostate adenocarcinoma 4	1.7
Colon NAT 4	2.4	Prostate NAT 5	1.1
Lung cancer 1	2.9	Prostate adenocarcinoma 6	2.6
Lung NAT 1	1.1	Prostate adenocarcinoma 7	3.3
Lung cancer 2	16.2	Prostate adenocarcinoma 8	0.6
Lung NAT 2	2.3	Prostate adenocarcinoma 9	6.5
Squamous cell carcinoma 3	4.8	Prostate NAT 10	1.4
Lung NAT 3	0.5	Kidney cancer 1	14.2
Metastatic melanoma 1	8.7	Kidney NAT 1	7.6
Melanoma 2	3.7	Kidney cancer 2	100.0
Melanoma 3	9.2	Kidney NAT 2	15.6
Metastatic melanoma 4	16.3	Kidney cancer 3	38.7
Metastatic melanoma 5	20.2	Kidney NAT 3	6.5
Bladder cancer 1	1.3	Kidney cancer 4	11.8
Bladder NAT 1	0.0	Kidney NAT 4	6.9
Bladder cancer 2	3.9		

CNS_neurodegeneration_v1.0 Summary: Ag5627 Two experiments with same probe-primer sets are in good agreements. This panel confirms the expression of this gene at low levels in the brain in an independent group of individuals. This gene is found to be upregulated in the temporal cortex of Alzheimer's disease patients. Therefore, therapeutic

5   modulation of the expression or function of this gene may decrease neuronal death and be of use in the treatment of this disease.

Panel 4.1D Summary: Ag5627 Highest expression of this gene is detected in kidney. Moderate to low levels of expression of this gene is also seen in activated naive and memory T cells, IL-2 treated NK cells, IFN gamma activated HUVEC cells, cytokine

10   activated bronchial epithelial cells, astrocytes, resting and activated small airway epithelial cells, coronery artery SMC cells, basophils, keratinocytes, mucoepidermoid NCI-H292 cells, lung and dermal fibroblast, liver cirrhosis sample and normal tissues such as colon, lung, and thymus. Therefore, therapeutic modulation of this gene or its protein product through the use of small molecule drug may be useful in the treatment of autoimmune and

15   inflammatory diseases such as asthma, allergies, inflammatory bowel disease, lupus erythematosus, psoriasis, rheumatoid arthritis, and osteoarthritis.

Panel 5 Islet Summary: Ag5627 Two experiments with same probe and primer sets are in good agreements. Highest expression of this gene is detected in placenta of diabetic and nondiabetic patients (CTs=26.4-26.7). Moderate to high levels of expression of

20   this gene is also seen in liver HepG2 cell line, adipose, small intestine and kidney. This gene codes for a homolog of Serine palmitoyltransferase 2. Serine palmitoyltransferase catalyzes the first, rate limiting step in de novo ceramide biosynthesis. C2-ceramide inhibits GLUT4 translocation by inhibiting Akt phosphorylation and activation in 3T3-L1 adipocytes, independently of effects on IRS-1 (Summers et al., 1998, Mol Cell Biol

25   18:5457-64, PMID: 9710629). Ceramide downregulates PDE3B and induces lipolysis in 3T3-L1 cells. Ceramide effects are reversed by troglitazone (Mei et al., 2002, Diabetes 51: 631-7, PMID: 11872660). Palmitate-induced insulin resistance involves elevation of de novo ceramide synthesis in C2C12 myotubes (Schmitz-Peiffer et al., 1999, J Biol Chem 274:24202, PMID: 10446195). Therefore, inhibition of the novel serine

30   palmitoyltransferase through the use of small molecule drug may be beneficial in the treatment of diabetes.

general oncology screening panel_v_2.4 Summary: Ag5627 Highest expression of this gene is detected in kidney cancer (CT=27.5). Moderate to high expression of this

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gene is also seen in normal and cancer samples derived from colon, lung, bladder, prostate and kidney. Moderate levels of expression of this gene is also seen in melanoma and metastatic melanoma samples. Expression of this gene is strongly associated with kidney, lung and bladder cancers as compared to the corresponding normal tissues. Therefore,

5   expression of this gene may be used as diagnostic marker for detection of these cancers and also, therapeutic modulation of this gene or its protein product may be useful in the treatment of melanoma, colon, lung, bladder, prostate and kidney cancers.

AA. CG148888-01: GALNAC 4-SULFOTRANSFERASE.

Expression of gene CG148888-01 was assessed using the primer-probe set Ag6854,
10   described in Table AAA. Results of the RTQ-PCR runs are shown in Table AAB. Please note that CG148888-01 represents a full-length physical clone.

Table AAA. Probe Name Ag6854

Primers	Sequenes	Length	Start Position	SEQ ID No
Forward	5'-accccagagccgcctggt-3'	18	369	345
Probe	TET-5'-cttggcctgatgttgaactttattcctg gcacc-3'-TAMRA	33	408	346
Reverse	5'-cagcctgcaggaccctacg-3'	19	458	347

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Table AAB. General screening panel v1.6

Tissue Name	Rel. Exp.(%) Ag6854, Run 278020603	issue Name	Rel. Exp.(%) Ag6854, Run 278020603
Adipose	0.0	Renal ca. TK-10	0.0
Melanoma* Hs688(A).T	0.0	Bladder	0.1
Melanoma* Hs688(B).T	0.2	Gastric ca. (liver met.) NCI-N87	0.0
Melanoma* M14	0.0	Gastric ca. KATO III	0.0
Melanoma* LOXIMVI	0.0	Colon ca. SW-948	0.0
Melanoma* SK-MEL-5	0.3	Colon ca. SW480	0.1
Squamous cell carcinoma SCC-4	0.1	Colon ca.* (SW480 met) SW620	0.0
Testis Pool	0.2	Colon ca. HT29	0.0
Prostate ca.* (bone met) PC-3	0.0	Colon ca. HCT-116	0.0
Prostate Pool	0.0	Colon ca. CaCo-2	0.0

Placenta	0.0	Colon cancer tissue	0.1
Uterus Pool	0.0	Colon ca. SW1116	0.0
Ovarian ca. OVCAR-3	0.0	Colon ca. Colo-205	0.0
Ovarian ca. SK-OV-3	0.0	Colon ca. SW-48	0.0
Ovarian ca. OVCAR-4	0.0	Colon Pool	0.2
Ovarian ca. OVCAR-5	0.1	Small Intestine Pool	0.1
Ovarian ca. IGROV-1	0.0	Stomach Pool	0.3
Ovarian ca. OVCAR-8	0.0	Bone Marrow Pool	0.1
Ovary	0.2	Fetal Heart	0.3
Breast ca. MCF-7	0.7	Heart Pool	0.0
Breast ca. MDA-MB-231	0.0	Lymph Node Pool	0.5
Breast ca. BT 549	0.0	Fetal Skeletal Muscle	0.0
Breast ca. T47D	0.0	Skeletal Muscle Pool	0.0
Breast ca. MDA-N	0.0	Spleen Pool	0.6
Breast Pool	0.2	Thymus Pool	0.5
Trachea	0.3	CNS cancer (glio/astro) U87-MG	0.0
Lung	0.2	CNS cancer (glio/astro) U-118-MG	0.0
Fetal Lung	0.0	CNS cancer (neuro;met) SK-N-AS	2.2
Lung ca. NCI-N417	0.0	CNS cancer (astro) SF-539	0.0
Lung ca. LX-1	0.0	CNS cancer (astro) SNB-75	0.7
Lung ca. NCI-H146	0.0	CNS cancer (glio) SNB-19	0.0
Lung ca. SHP-77	100.0	CNS cancer (glio) SF-295	0.1
Lung ca. A549	0.0	Brain (Amygdala) Pool	3.7
Lung ca. NCI-H526	0.4	Brain (cerebellum)	8.8
Lung ca. NCI-H23	0.2	Brain (fetal)	16.2
Lung ca. NCI-H460	0.1	Brain (Hippocampus) Pool	3.6
Lung ca. HOP-62	0.0	Cerebral Cortex Pool	3.7
Lung ca. NCI-H522	1.4	Brain (Substantia nigra) Pool	4.6
Liver	0.0	Brain (Thalamus) Pool	5.0
Fetal Liver	0.0	Brain (whole)	4.5
Liver ca. HepG2	0.0	Spinal Cord Pool	4.7
Kidney Pool	0.0	Adrenal Gland	0.2
Fetal Kidney	0.0	Pituitary gland Pool	8.0
Renal ca. 786-0	0.0	Salivary Gland	0.0
Renal ca. A498	0.0	Thyroid (female)	0.2
Renal ca. ACHN	0.0	Pancreatic ca. CAPAN2	0.1
Renal ca. UO-31	0.0	Pancreas Pool	0.2

General_screening_panel_v1.6 Summary: Ag6854 Highest expression of this gene is seen in a lung cancer cell line (CT=27.8). Thus, expression of this gene could be used to differentiate between this sample and other samples on this panel and as a marker to

detect the presence of lung cancer. Furthermore, therapeutic modulation of the expression or function of this gene may be effective in the treatment of lung cancer.

This gene is also expressed at moderate to low levels in the CNS, including the hippocampus, thalamus, substantia nigra, amygdala, cerebellum and cerebral cortex.

5   Therefore, therapeutic modulation of the expression or function of this gene may be useful in the treatment of neurological disorders, such as Alzheimer's disease, Parkinson's disease, schizophrenia, multiple sclerosis, stroke and epilepsy.

AB. CG149008-01: NOVEL SODIUM/HYDROGEN EXCHANGER FAMILY MEMBER.

10   Expression of gene CG149008-01 was assessed using the primer-probe set Ag5630, described in Table ABA. Results of the RTQ-PCR runs are shown in Tables ABB, ABC, ABD and ABE.

Table ABA. Probe Name Ag5630

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Primers	Sequencs	Length	Start Position	SEQ ID No
Forward	5'-tattttctgggtcaggctgat-3'	21	770	348
Probe	TET-5'-tctctaaactcaacatgacagacagttttg-3'-TAMRA	30	795	349
Reverse	5'-cagatattagggagccaaacg-3'	21	825	350

Table ABB. CNS_neurodegeneration_v1.0

20

Tissue Name	Rel. Exp.(%) Ag5630, Run 246956911	issue Name	Rel. Exp.(%) Ag5630, Run 246956911
AD 1 Hippo	9.3	Control (Path) 3 Temporal Ctx	9.3
AD 2 Hippo	31.4	Control (Path) 4 Temporal Ctx	14.5
AD 3 Hippo	5.5	AD 1 Occipital Ctx	7.5
AD 4 Hippo	8.4	AD 2 Occipital Ctx (Missing)	0.0
AD 5 hippo	62.0	AD 3 Occipital Ctx	4.5
AD 6 Hippo	46.0	AD 4 Occipital Ctx	18.9
Control 2 Hippo	31.4	AD 5 Occipital Ctx	13.9
Control 4 Hippo	15.9	AD 6 Occipital Ctx	46.3
Control (Path) 3 Hippo	10.4	Control 1 Occipital Ctx	3.8

AD 1 Temporal Ctx	12.0	Control 2 Occipital Ctx	61.6
AD 2 Temporal Ctx	41.8	Control 3 Occipital Ctx	6.1
AD 3 Temporal Ctx	2.3	Control 4 Occipital Ctx	13.2
AD 4 Temporal Ctx	25.7	Control (Path) 1 Occipital Ctx	62.0
AD 5 Inf Temporal Ctx	100.0	Control (Path) 2 Occipital Ctx	10.5
AD 5 SupTemporal Ctx	48.6	Control (Path) 3 Occipital Ctx	8.4
AD 6 Inf Temporal Ctx	36.9	Control (Path) 4 Occipital Ctx	11.8
AD 6 Sup Temporal Ctx	45.7	Control 1 Parietal Ctx	10.4
Control 1 Temporal Ctx	14.3	Control 2 Parietal Ctx	49.0
Control 2 Temporal Ctx	48.6	Control 3 Parietal Ctx	20.3
Control 3 Temporal Ctx	12.8	Control (Path) 1 Parietal Ctx	44.1
Control 4 Temporal Ctx	14.1	Control (Path) 2 Parietal Ctx	22.7
Control (Path) 1 Temporal Ctx	52.5	Control (Path) 3 Parietal Ctx	8.2
Control (Path) 2 Temporal Ctx	33.9	Control (Path) 4 Parietal Ctx	35.1

Table ABC. General screening panel v1.5

5

Tissue Name	Rel. Exp.(%) Ag5630, Run 245065625	issue Name	Rel. Exp.(%) Ag5630, Run 245065625
Adipose	4.2	Renal ca. TK-10	32.8
Melanoma* Hs688(A).T	21.9	Bladder	9.5
Melanoma* Hs688(B).T	19.2	Gastric ca. (liver met.) NCI-N87	100.0
Melanoma* M14	41.2	Gastric ca. KATO III	52.1
Melanoma* LOXIMVI	25.2	Colon ca. SW-948	5.1
Melanoma* SK-MEL-5	20.0	Colon ca. SW480	27.2
Squamous cell carcinoma SCC-4	8.4	Colon ca.* (SW480 met) SW620	22.2
Testis Pool	9.1	Colon ca. HT29	10.5
Prostate ca.* (bone met) PC-3	5.8	Colon ca. HCT-116	15.6
Prostate Pool	3.0	Colon ca. CaCo-2	25.9
Placenta	16.7	Colon cancer tissue	12.9
Uterus Pool	4.3	Colon ca. SW1116	3.4
Ovarian ca. OVCAR-3	35.6	Colon ca. Colo-205	19.8
Ovarian ca. SK-OV-3	15.4	Colon ca. SW-48	12.6
Ovarian ca. OVCAR-4	9.5	Colon Pool	6.4
Ovarian ca. OVCAR-5	44.8	Small Intestine Pool	4.0
Ovarian ca. IGROV-1	13.9	Stomach Pool	3.7
Ovarian ca. OVCAR-8	8.0	Bone Marrow Pool	2.9
Ovary	3.8	Fetal Heart	4.1
Breast ca. MCF-7	14.9	Heart Pool	3.3
Breast ca. MDA-MB-231	25.2	Lymph Node Pool	6.8

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Breast ca. BT 549	32.1	Fetal Skeletal Muscle	2.5
Breast ca. T47D	18.7	Skeletal Muscle Pool	15.6
Breast ca. MDA-N	9.3	Spleen Pool	5.4
Breast Pool	1.7	Thymus Pool	7.6
Trachea	18.4	CNS cancer (glio/astro) U87-MG	74.2
Lung	1.7	CNS cancer (glio/astro) U-118-MG	34.4
Fetal Lung	9.2	CNS cancer (neuro;met) SK-N-AS	8.5
Lung ca. NCI-N417	4.8	CNS cancer (astro) SF-539	11.9
Lung ca. LX-1	24.1	CNS cancer (astro) SNB-75	43.2
Lung ca. NCI-H146	3.6	CNS cancer (glio) SNB-19	12.9
Lung ca. SHP-77	14.0	CNS cancer (glio) SF-295	30.8
Lung ca. A549	35.4	Brain (Amygdala) Pool	4.9
Lung ca. NCI-H526	3.5	Brain (cerebellum)	23.7
Lung ca. NCI-H23	23.5	Brain (fetal)	6.5
Lung ca. NCI-H460	6.7	Brain (Hippocampus) Pool	7.5
Lung ca. HOP-62	7.6	Cerebral Cortex Pool	5.3
Lung ca. NCI-H522	8.5	Brain (Substantia nigra) Pool	4.3
Liver	4.2	Brain (Thalamus) Pool	7.4
Fetal Liver	15.8	Brain (whole)	5.4
Liver ca. HepG2	5.7	Spinal Cord Pool	6.4
Kidney Pool	7.7	Adrenal Gland	24.1
Fetal Kidney	5.0	Pituitary gland Pool	3.1
Renal ca. 786-0	19.9	Salivary Gland	13.2
Renal ca. A498	14.3	Thyroid (female)	8.1
Renal ca. ACHN	8.9	Pancreatic ca. CAPAN2	26.1
Renal ca. UO-31	32.1	Pancreas Pool	9.3

Table ABD. Panel 4.1D

5

Tissue Name	Rel. Exp.(% Ag5630, Run 246490808	Tissue Name	Rel. Exp.(%) Ag5630, Run 246490808
Secondary Th1 act	52.9	HUVEC IL-1beta	21.9
Secondary Th2 act	86.5	HUVEC IFN gamma	20.2
Secondary Tr1 act	14.5	HUVEC TNF alpha + IFN gamma	6.7
Secondary Th1 rest	2.2	HUVEC TNF alpha + IL4	4.6
Secondary Th2 rest	1.7	HUVEC IL-11	12.6
Secondary Tr1 rest	0.0	Lung Microvascular EC none	31.6
Primary Th1 act	0.8	Lung Microvascular EC TNFalpha + IL-1beta	9.4
Primary Th2 act	42.6	Microvascular Dermal EC none	0.7

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Primary Tr1 act	35.4	Microsvastilar Dermal EC TNFalpha + IL-1beta	7.2
Primary Th1 rest	1.9	Bronchial epithelium TNFalpha + IL1beta	4.2
Primary Th2 rest	3.4	Small airway epithelium none	4.5
Primary Tr1 rest	0.3	Small airway epithelium TNFalpha + IL-1beta	29.1
CD45RA CD4 lymphocyte act	30.6	Coronery artery SMC rest	9.9
CD45RO CD4 lymphocyte act	49.3	Coronery artery SMC TNFalpha + IL-1beta	13.3
CD8 lymphocyte act	4.6	Astrocytes rest	2.6
Secondary CD8 lymphocyte rest	29.9	Astrocytes TNFalpha + IL-1beta	4.2
Secondary CD8 lymphocyte act	6.6	KU-812 (Basophil) rest	4.9
CD4 lymphocyte none	0.0	KU-812 (Basophil) PMA/ionomycin	11.9
2ry Th1/Th2/Tr1_anti-CD95 CH11	2.5	CCD1106 (Keratinocytes) none	28.3
LAK cells rest	11.1	CCD1106 (Keratinocytes) TNFalpha + IL-1beta	18.6
LAK cells IL-2	9.7	Liver cirrhosis	4.6
LAK cells IL-2+IL-12	2.3	NCI-H292 none	46.3
LAK cells IL-2+IFN gamma	17.3	NCI-H292 IL-4	46.0
LAK cells IL-2+ IL-18	9.5	NCI-H292 IL-9	69.3
LAK cells PMA/ionomycin	36.3	NCI-H292 IL-13	59.0
NK Cells IL-2 rest	17.0	NCI-H292 IFN gamma	33.9
Two Way MLR 3 day	9.4	HPAEC none	12.9
Two Way MLR 5 day	1.0	HPAEC TNF alpha + IL-1 beta	70.2
Two Way MLR 7 day	7.0	Lung fibroblast none	14.2
PBMC rest	0.9	Lung fibroblast TNF alpha + IL-1 beta	20.0
PBMC PWM	9.9	Lung fibroblast IL-4	12.4
PBMC PHA-L	8.4	Lung fibroblast IL-9	4.8
Ramos (B cell) none	1.4	Lung fibroblast IL-13	2.7
Ramos (B cell) ionomycin	28.5	Lung fibroblast IFN gamma	27.7
B lymphocytes PWM	19.6	Dermal fibroblast CCD1070 rest	33.9
B lymphocytes CD40L and IL-4	28.1	Dermal fibroblast CCD1070 TNF alpha	62.4
EOL-1 dbcAMP	3.8	Dermal fibroblast CCD1070 IL-1 beta	18.3
EOL-1 dbcAMP PMA/ionomycin	0.4	Dermal fibroblast IFN gamma	19.3
Dendritic cells none	9.2	Dermal fibroblast IL-4	37.4
Dendritic cells LPS	3.2	Dermal Fibroblasts rest	15.8
Dendritic cells anti-CD40	3.8	Neutrophils TNFa+LPS	37.6
Monocytes rest	0.0	Neutrophils rest	41.2
Monocytes LPS	100.0	Colon	1.5

Macrophages rest	6.0	Lung	0.9
Macrophages LPS	10.6	Thymus	2.4
HUVEC none	12.6	Kidney	17.2
HUVEC starved	21.5		

Table ABE. Panel 5 Islet

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Tissue Name	Rel. Exp.(% Ag5630, Run 279370866	Tissue Name	Rel. Exp.(%) Ag5630, Run 279370866
97457_Patient-02go_adipose	15.5	94709_Donor 2 AM - A_adipose	26.6
97476_Patient-07sk_skeletal muscle	0.0	94710_Donor 2 AM - B_adipose	21.0
97477_Patient-07ut_uterus	5.0	94711_Donor 2 AM - C_adipose	16.7
97478_Patient-07pl_placenta	9.3	94712_Donor 2 AD - A_adipose	55.9
99167_Bayer Patient 1	100.0	94713_Donor 2 AD - B_adipose	74.7
97482_Patient-08ut_uterus	11.0	94714_Donor 2 AD - C_adipose	54.7
97483_Patient-08pl_placenta	7.9	94742_Donor 3 U - A_Mesenchymal Stem Cells	5.7
97486_Patient-09sk_skeletal muscle	9.9	94743_Donor 3 U - B_Mesenchymal Stem Cells	8.0
97487_Patient-09ut_uterus	4.1	94730_Donor 3 AM - A_adipose	8.3
97488_Patient-09pl_placenta	10.3	94731_Donor 3 AM - B_adipose	14.3
97492_Patient-10ut_uterus	10.2	94732_Donor 3 AM - C_adipose	11.3
97493_Patient-10pl_placenta	20.9	94733_Donor 3 AD - A_adipose	30.1
97495_Patient-11go_adipose	5.8	94734_Donor 3 AD - B_adipose	22.5
97496_Patient-11sk_skeletal muscle	4.4	94735_Donor 3 AD - C_adipose	7.5
97497_Patient-11ut_uterus	13.5	77138_Liver_HepG2untreated	2.5
97498_Patient-11pl_placenta	3.4	73556_Heart_Cardiac stromal cells (primary)	2.7
97500_Patient-12go_adipose	37.1	81735_Small Intestine	12.6
97501_Patient-12sk_skeletal muscle	20.2	72409_Kidney_Proximal Convoluted Tubule	28.1
97502_Patient-12ut_uterus	22.8	82685_Small intestine_Duodenum	24.0
97503_Patient-12pl_placenta	13.1	90650_Adrenal_Adrenocortical adenoma	7.3
94721_Donor 2 U - A_Mesenchymal Stem Cells	87.7	72410_Kidney_HRCE	33.0
94722_Donor 2 U - B_Mesenchymal Stem Cells	75.8	72411_Kidney_HRE	10.4
94723_Donor 2 U - C_Mesenchymal Stem Cells	77.9	73139_Uterus_Uterine smooth muscle cells	11.8

CNS_neurodegeneration_v1.0 Summary: Ag5630 This panel confirms the expression of this gene at low levels in the brains of an independent group of individuals.

5   However, no differential expression of this gene was detected between Alzheimer's diseased postmortem brains and those of non-demented controls in this experiment. Please see Panel 1.5 for a discussion of this gene in treatment of central nervous system disorders.

General_screening_panel_v1.5 Summary: Ag5630 Higest expression of this gene is detected in a gastric cancer NCI-N87 cell line (CT=27.6). Moderate levels of

10   expression of this gene is also seen in cluster of cancer cell lines derived from pancreatic, gastric, colon, lung, liver, renal, breast, ovarian, prostate, squamous cell carcinoma, melanoma and brain cancers. Thus, expression of this gene could be used as a marker to detect the presence of these cancers. Furthermore, therapeutic modulation of the expression or function of this gene may be effective in the treatment of pancreatic, gastric, colon, lung,

15   liver, renal, breast, ovarian, prostate, squamous cell carcinoma, melanoma and brain cancers.

Among tissues with metabolic or endocrine function, this gene is expressed at moderate levels in pancreas, adipose, adrenal gland, thyroid, pituitary gland, skeletal muscle, heart, liver and the gastrointestinal tract. Therefore, therapeutic modulation of the

20   activity of this gene may prove useful in the treatment of endocrine/metabolically related diseases, such as obesity and diabetes.

In addition, this gene is expressed at moderate levels in all regions of the central nervous system examined, including amygdala, hippocampus, substantia nigra, thalamus, cerebellum, cerebral cortex, and spinal cord. Therefore, therapeutic modulation of this gene

25   product may be useful in the treatment of central nervous system disorders such as Alzheimer's disease, Parkinson's disease, epilepsy, multiple sclerosis, schizophrenia and depression.

Panel 4.1D Summary: Ag5630 Higest expression of this gene is detected in LPS treated monocytes (CT=29.7). Interestingly, this gene is expressed at much higher levels in

30   LPS activated when compared to resting monocytes (CT=40). This observation suggests that expression of this gene can be used to distinguish actvated from resting monocytes. In addition, upon activation monocytes contribute to the innate and specific immunity by migrating to the site of tissue injury and releasing inflammatory cytokines. This release

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contributes to the inflammation process. Therefore, modulation of the expression of the protein encoded by this gene may prevent the recruitment of monocytes and the initiation of the inflammatory process.

In addition, this gene is also expressed at moderate to low levels in activated polarized T cells, naive and memory T cells, resting and activated LAK cells, resting IL-2 treated NK cells, two way MLR, activated PBMC cells and B lymphocytes, dendritic cells, macrophage, different endothelial cells, bronchial and small airway epithelium, astrocytes, basophils, keratinocytes, mucoepidermoid cells, lung and dermal fibroblasts, neutrophils and kidney. Therefore, modulation of the gene product with a functional therapeutic may lead to the alteration of functions associated with these cell types and lead to improvement of the symptoms of patients suffering from autoimmune and inflammatory diseases such as asthma, allergies, inflammatory bowel disease, lupus erythematosus, psoriasis, rheumatoid arthritis, and osteoarthritis.

Panel 5 Islet Summary: Ag5630 Higest expression of this gene is detected in beta islet cells (CT=26.7). In addition, this gene shows widespread expression in this panel, with moderate to low expressions in adipose, placenta, uterus, skeletal muscle, kidney, and small intestine samples. Therefore, therapeutic modulation of this gene may be useful in the treatment of metabolic/endocrine disorders including, obesity, Type I and II diabetes.

AC. CG149350-01 and CG149350-02: Vacuolar ATP synthase subunit F.

Expression of gene CG149350-01 and CG149350-02 was assessed using the primer-probe set Ag7581, described in Table ACA. Results of the RTQ-PCR runs are shown in Table ACB. Please note that CG149350-02 represents a full-length physical clone of the CG149350-01 gene, validating the prediction of the gene sequence.

Table ACA. Probe Name Ag7581

Primers		Length	Start Position	SEQ ID No
Forward	5'-aagaactgccaccccaatt-3'	19	88	351
Probe	TET-5'-cattgatggtcgtatccttctccacca-3'-TAMRA	27	113	352
Reverse	5'-aaattgccggaaagtgtctt-3'	20	146	353

Table ACB. CNS_neurodegeneration_v1.0

Tissue Name	Rel. Exp.(%) Ag7581, Run 308752174	issue Name	Rel. Exp.(%) Ag7581, Run 308752174
AD 1 Hippo	19.9	Control (Path) 3 Temporal Ctx	7.3
AD 2 Hippo	21.3	Control (Path) 4 Temporal Ctx	62.9
AD 3 Hippo	14.9	AD 1 Occipital Ctx	19.1
AD 4 Hippo	6.4	AD 2 Occipital Ctx (Missing)	0.0
AD 5 hippo	65.5	AD 3 Occipital Ctx	22.4
AD 6 Hippo	44.4	AD 4 Occipital Ctx	32.3
Control 2 Hippo	21.9	AD 5 Occipital Ctx	4.4
Control 4 Hippo	30.6	AD 6 Occipital Ctx	20.2
Control (Path) 3 Hippo	10.7	Control 1 Occipital Ctx	3.0
AD 1 Temporal Ctx	23.0	Control 2 Occipital Ctx	35.6
AD 2 Temporal Ctx	27.5	Control 3 Occipital Ctx	53.2
AD 3 Temporal Ctx	19.8	Control 4 Occipital Ctx	6.8
AD 4 Temporal Ctx	21.3	Control (Path) 1 Occipital Ctx	70.7
AD 5 Inf Temporal Ctx	46.3	Control (Path) 2 Occipital Ctx	17.9
AD 5 SupTemporal Ctx	55.9	Control (Path) 3 Occipital Ctx	4.2
AD 6 Inf Temporal Ctx	52.9	Control (Path) 4 Occipital Ctx	32.5
AD 6 Sup Temporal Ctx	47.3	Control 1 Parietal Ctx	8.7
Control 1 Temporal Ctx	23.5	Control 2 Parietal Ctx	56.3
Control 2 Temporal Ctx	28.9	Control 3 Parietal Ctx	32.5
Control 3 Temporal Ctx	22.2	Control (Path) 1 Parietal Ctx	100.0
Control 4 Temporal Ctx	9.1	Control (Path) 2 Parietal Ctx	38.4
Control (Path) 1 Temporal Ctx	45.7	Control (Path) 3 Parietal Ctx	17.6
Control (Path) 2 Temporal Ctx	62.0	Control (Path) 4 Parietal Ctx	64.2

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CNS_neurodegeneration_v1.0 Summary: Ag7581 No differential expression of this gene was detected between Alzheimer's diseased postmortem brains and those of non-demented controls in this experiment. However, this panel confirms the expression of this gene at low levels in the brains of an independent group of individuals. Therefore,

10   therapeutic modulation of this gene product may be useful in the treatment of central nervous system disorders such as Parkinson's disease, epilepsy, multiple sclerosis, schizophrenia and depression.

AD. CG149536-01: PROTEIN-TYROSINE PHOSPHATASE, NON-RECEPTOR TYPE 2.

Expression of gene CG149536-01 was assessed using the primer-probe sets Ag5255 and Ag6844, described in Tables ADA and ADB. Results of the RTQ-PCR runs are shown in Tables ADC, ADD and ADE.

Table ADA. Probe Name Ag5255

Primers		Length	Start Position	SEQ ID No
Forward	5'-cttatggtttggcagcagaa-3'	20	355	354
Probe	TET-5'-ccaaagcagttgtcatgctgaaccgc-3'-TAMRA	26	377	355
Reverse	5'-tggtttcaccactcgattct-3'	20	414	356

Table ADB. Probe Name Ag6844

Primers		Length	Start Position	SEQ ID No
Forward	5'-agagaatcgagtggtgaaacc-3'	21	412	357
Probe	TET-5'-actacctggccagattttggagtccc-3'-TAMRA	26	457	358
Reverse	5'-aggagccagattctctcacttta-3'	23	516	359

Table ADC. CNS_neurodegeneration_v1.0

Tissue Name	Rel. Exp.(%) Ag5255, Run 229929883	issue Name	Rel. Exp.(%) Ag5255, Run 229929883
AD 1 Hippo	28.9	Control (Path) 3 Temporal Ctx	21.0
AD 2 Hippo	42.3	Control (Path) 4 Temporal Ctx	38.7
AD 3 Hippo	42.0	AD 1 Occipital Ctx	45.4
AD 4 Hippo	5.9	AD 2 Occipital Ctx (Missing)	0.0
AD 5 hippo	92.7	AD 3 Occipital Ctx	36.9
AD 6 Hippo	29.7	AD 4 Occipital Ctx	23.5
Control 2 Hippo	52.5	AD 5 Occipital Ctx	13.6
Control 4 Hippo	22.4	AD 6 Occipital Ctx	47.6
Control (Path) 3 Hippo	17.9	Control 1 Occipital Ctx	3.2

AD 1 Temporal Ctx	39.5	Control 2 Occipital Ctx	57.4
AD 2 Temporal Ctx	56.3	Control 3 Occipital Ctx	31.2
AD 3 Temporal Ctx	23.3	Control 4 Occipital Ctx	5.0
AD 4 Temporal Ctx	10.9	Control (Path) 1 Occipital Ctx	99.3
AD 5 Inf Temporal Ctx	44.8	Control (Path) 2 Occipital Ctx	40.3
AD 5 SupTemporal Ctx	53.2	Control (Path) 3 Occipital Ctx	0.0
AD 6 Inf Temporal Ctx	68.8	Control (Path) 4 Occipital Ctx	24.0
AD 6 Sup Temporal Ctx	100.0	Control 1 Parietal Ctx	20.6
Control 1 Temporal Ctx	13.4	Control 2 Parietal Ctx	68.3
Control 2 Temporal Ctx	34.4	Control 3 Parietal Ctx	29.5
Control 3 Temporal Ctx	84.1	Control (Path) 1 Parietal Ctx	46.3
Control 4 Temporal Ctx	18.4	Control (Path) 2 Parietal Ctx	31.2
Control (Path) 1 Temporal Ctx	41.2	Control (Path) 3 Parietal Ctx	6.9
Control (Path) 2 Temporal Ctx	58.6	Control (Path) 4 Parietal Ctx	45.1

Table ADD. General screening panel v1.5

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Tissue Name	Rel. Exp.(%) Ag5255, Run 230218532	issue Name	Rel. Exp.(%) Ag5255, Run 230218532
Adipose	6.4	Renal ca. TK-10	18.8
Melanoma* Hs688(A).T	9.5	Bladder	10.8
Melanoma* Hs688(B).T	8.7	Gastric ca. (liver met.) NCI-N87	50.3
Melanoma* M14	19.1	Gastric ca. KATO III	60.3
Melanoma* LOXIMVI	25.5	Colon ca. SW-948	5.8
Melanoma* SK-MEL-5	18.8	Colon ca. SW480	100.0
Squamous cell carcinoma SCC-4	24.0	Colon ca.* (SW480 met) SW620	23.3
Testis Pool	2.2	Colon ca. HT29	19.2
Prostate ca.* (bone met) PC-3	33.9	Colon ca. HCT-116	46.7
Prostate Pool	4.1	Colon ca. CaCo-2	49.3
Placenta	1.9	Colon cancer tissue	5.7
Uterus Pool	2.3	Colon ca. SW1116	3.5
Ovarian ca. OVCAR-3	19.6	Colon ca. Colo-205	3.3
Ovarian ca. SK-OV-3	55.5	Colon ca. SW-48	0.5
Ovarian ca. OVCAR-4	8.5	Colon Pool	5.9
Ovarian ca. OVCAR-5	44.4	Small Intestine Pool	5.7
Ovarian ca. IGROV-1	5.7	Stomach Pool	3.2
Ovarian ca. OVCAR-8	7.8	Bone Marrow Pool	2.8
Ovary	8.0	Fetal Heart	3.7
Breast ca. MCF-7	38.2	Heart Pool	0.7
Breast ca. MDA-MB-231	13.4	Lymph Node Pool	4.1

Breast ca. BT 549	51.8	Fetal Skeletal Muscle	5.8
Breast ca. T47D	5.4	Skeletal Muscle Pool	2.6
Breast ca. MDA-N	7.0	Spleen Pool	0.4
Breast Pool	9.0	Thymus Pool	19.2
Trachea	1.0	CNS cancer (glio/astro) U87-MG	26.4
Lung	5.7	CNS cancer (glio/astro) U-118-MG	33.2
Fetal Lung	17.1	CNS cancer (neuro;met) SK-N-AS	18.9
Lung ca. NCI-N417	1.0	CNS cancer (astro) SF-539	17.1
Lung ca. LX-1	12.6	CNS cancer (astro) SNB-75	12.2
Lung ca. NCI-H146	16.6	CNS cancer (glio) SNB-19	6.4
Lung ca. SHP-77	34.6	CNS cancer (glio) SF-295	16.0
Lung ca. A549	15.1	Brain (Amygdala) Pool	4.0
Lung ca. NCI-H526	6.7	Brain (cerebellum)	33.2
Lung ca. NCI-H23	33.0	Brain (fetal)	54.0
Lung ca. NCI-H460	7.2	Brain (Hippocampus) Pool	4.7
Lung ca. HOP-62	26.2	Cerebral Cortex Pool	5.3
Lung ca. NCI-H522	35.1	Brain (Substantia nigra) Pool	4.0
Liver	0.9	Brain (Thalamus) Pool	6.8
Fetal Liver	7.2	Brain (whole)	4.9
Liver ca. HepG2	9.7	Spinal Cord Pool	7.0
Kidney Pool	7.3	Adrenal Gland	2.4
Fetal Kidney	16.3	Pituitary gland Pool	2.1
Renal ca. 786-0	7.1	Salivary Gland	1.5
Renal ca. A498	2.2	Thyroid (female)	1.1
Renal ca. ACHN	9.2	Pancreatic ca. CAPAN2	66.4
Renal ca. UO-31	6.5	Pancreas Pool	7.2

Table ADE. Panel 4.1D

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Tissue Name	Rel. Exp.(%) g5255, Run 229851730	Rel. Exp.(%) Ag6844, Run 279029113	Tissue Name	Rel. Exp.(%) Ag5255, Run 229851730	Rel. Exp.(%) Ag6844, Run 279029113
Secondary Th1 act	39.0	38.7	HUVEC IL-1beta	39.8	9.6
Secondary Th2 act	46.7	55.9	HUVEC IFN gamma	12.5	15.9
Secondary Tr1 act	15.7	18.9	HUVEC TNF alpha + IFN gamma	21.0	8.4
Secondary Th1 rest	12.0	3.9	HUVEC TNF alpha + IL4	12.1	11.0
Secondary Th2 rest	0.0	5.3	HUVEC IL-11	13.6	4.4
Secondary Tr1 rest	0.0	9.2	Lung Microvascular EC none	25.2	18.4

Primary Th1 act	17.9	6.0	Lung Microvascular EC TNFalpha + IL-1beta	2.6	9.4
Primary Th2 act	15.0	33.7	Microvascular Dermal EC none	6.0	3.8
Primary Tr1 act	18.2	22.7	Microsvasular Dermal EC TNFalpha + IL-1beta	0.0	3.7
Primary Th1 rest	0.0	1.9	Bronchial epithelium TNFalpha + IL1beta	9.3	10.2
Primary Th2 rest	5.0	1.5	Small airway epithelium none	0.0	10.0
Primary Tr1 rest	0.0	0.0	Small airway epithelium TNFalpha + IL-1beta	37.1	14.1
CD45RA CD4 lymphocyte act	32.1	13.9	Coronery artery SMC rest	11.1	5.5
CD45RO CD4 lymphocyte act	58.6	42.9	Coronery artery SMC TNFalpha + IL-1beta	11.3	4.0
CD8 lymphocyte act	5.2	18.7	Astrocytes rest	0.0	1.1
Secondary CD8 lymphocyte rest	10.9	5.5	Astrocytes TNFalpha + IL-1beta	0.0	1.8
Secondary CD8 lymphocyte act	0.0	4.4	KU-812 (Basophil) rest	38.4	17.2
CD4 lymphocyte none	6.7	3.4	KU-812 (Basophil) PMA/ionomycin	33.2	38.7
2ry Th1/Th2/Tr1_anti-CD95 CH11	0.0	26.4	CCD1106 (Keratinocytes) none	76.3	40.1
LAK cells rest	19.1	14.7	CCD1106 (Keratinocytes) TNFalpha + IL-1beta	13.1	14.9
LAK cells IL-2	5.4	7.3	Liver cirrhosis	15.8	7.0
LAK cells IL-2+IL-12	7.9	1.0	NCI-H292 none	35.1	20.2
LAK cells IL-2+IFN gamma	16.2	7.7	NCI-H292 IL-4	45.4	25.5
LAK cells IL-2+ IL-18	5.1	8.0	NCI-H292 IL-9	60.7	31.2
LAK cells PMA/ionomycin	27.9	40.9	NCI-H292 IL-13	45.4	38.4
NK Cells IL-2 rest	27.9	40.3	NCI-H292 IFN gamma	26.2	16.7
Two Way MLR 3 day	18.2	27.0	HPAEC none	5.6	6.3
Two Way MLR 5 day	23.3	2.1	HPAEC TNF alpha + IL-1 beta	21.5	12.1
Two Way MLR 7 day	4.5	1.7	Lung fibroblast none	22.5	12.2
PBMC rest	3.2	5.4	Lung fibroblast TNF alpha + IL-1 beta	6.3	8.2
PBMC PWM	20.6	9.8	Lung fibroblast IL-4	16.0	13.5

PBMC PHA-L	21.6	12.1	Lung fibroblast IL-9	15.9	11.9
Ramos (B cell) none	40.3	4.8	Lung fibroblast IL-13	0.0	5.8
Ramos (B cell) ionomycin	31.6	17.7	Lung fibroblast IFN gamma	37.6	19.9
B lymphocytes PWM	26.6	6.0	Dermal fibroblast CCD1070 rest	32.3	17.2
B lymphocytes CD40L and IL-4	4.8	37.6	Dermal fibroblast CCD1070 TNF alpha	100.0	54.7
EOL-1 dbcAMP	62.9	74.2	Dermal fibroblast CCD1070 IL-1 beta	34.6	18.7
EOL-1 dbcAMP PMA/ionomycin	45.4	15.1	Dermal fibroblast IFN gamma	17.1	12.7
Dendritic cells none	33.7	57.0	Dermal fibroblast IL-4	5.3	15.0
Dendritic cells LPS	21.0	15.2	Dermal Fibroblasts rest	0.0	6.9
Dendritic cells anti-CD40	10.2	7.3	Neutrophils TNFa+LPS	0.0	2.7
Monocytes rest	4.3	32.1	Neutrophils rest	5.6	6.1
Monocytes LPS	69.7	100.0	Colon	0.0	0.9
Macrophages rest	17.0	3.8	Lung	0.0	1.7
Macrophages LPS	0.0	9.3	Thymus	15.2	18.2
HUVEC none	5.9	28.7	Kidney	6.3	8.7
HUVEC starved	28.1	8.5			

AI_comprehensive panel_v1.0 Summary: Ag5255 Expression of this gene is low/undetectable (CTs > 35) across all of the samples on this panel.

CNS_neurodegeneration_v1.0 Summary: Ag5255 This panel confirms the expression of this gene at low levels in the brains of an independent group of individuals. However, no differential expression of this gene was detected between Alzheimer's diseased postmortem brains and those of non-demented controls in this experiment. Please see Panel 1.5 for a discussion of this gene in treatment of central nervous system disorders.

General_screening_panel_v1.5 Summary: Ag5255 Highest expression of this gene is detected in a colon cancer SW480 cell line (CT=31.6). Moderate to low levels of expression of this gene is also seen in cluster of cancer cell lines derived from pancreatic, gastric, colon, lung, liver, renal, breast, ovarian, prostate, squamous cell carcinoma, melanoma and brain cancers. Thus, expression of this gene could be used as a marker to detect the presence of these cancers. Furthermore, therapeutic modulation of the expression or function of this gene may be effective in the treatment of pancreatic, gastric, colon, lung,

liver, renal, breast, ovarian, prostate, squamous cell carcinoma, melanoma and brain cancers.

In addition, this gene is expressed at moderate levels in cerebellum and fetal brain. Therefore, therapeutic modulation of this gene product may be useful in the treatment of central nervous system disorders such ataxia and autism.

Panel 4.1D Summary: Ag5255/Ag6844 Two experiments with different probe and primer sets are in good agreement. The highest expression of this gene is detected in TNF alpha activated dermal fibroblast and LPS activated monocytes (CTs=32.7-32.9). Moderate to low levels of expression of this gene is detected in activated polarized T cells, naive and memory T cells, PMA/ionomycin activated LAK cells, resting IL-2 treated NK cells, eosinophils, resting dendritic cells, activated basophils, resting keratinocyte, and activated mucoepidermoid NCI-H292 cells. Therefore, therapeutic modulation of this gene or its protein product may be useful in the treatment of autoimmune and inflammatory diseases such as asthma, allergies, inflammatory bowel disease, lupus erythematosus, psoriasis, rheumatoid arthritis, and osteoarthritis.

AE. CG149964-01: Brain mitochondrial carrier protein-1.

Expression of gene CG149964-01 was assessed using the primer-probe set Ag7056, described in Table AEA.

Table AEA. Probe Name Ag7056

Primers	Sequencs	Length	Start Position	SEQ ID No
Forward	5'-tgtggttccaactgctcag-3'	19	617	360
Probe	TET-5'-ctggtagctctactcctacaacgatgg cag-3'-TAMRA	30	640	361
Reverse	5'-agatccacatgtcccatcatt-3'	21	707	362

General_screening_panel_v1.6 Summary: Ag7056 Expression of this gene is low/undetectable in all samples on this panel (CTs>35).

AF. CG150799-01, CG150799-02 and CG150799-03: MASS1.

Expression of gene CG150799-01, CG150799-02 and CG150799-03 was assessed using the primer-probe sets Ag5242, Ag5243, Ag5244, Ag5245, Ag5247 and Ag5248,

described in Tables AFA, AFB and AFC. Results of the RTQ-PCR runs are shown in Tables AFD, AFE, AFF, AFG, AFH and AFI. Please note that probe-primer sets Ag5243 is specific for CG150799-02 and probe-primer sets Ag5244 and Ag5245 are specific for CG150799-03.

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Table AFA. Probe Name Ag5242

Primers		Length	Start Position	SEQ ID No
Forward	5'-acgaatcccatgtgacactt-3'	21	3624	363
Probe	TET-5'-cccttcattataaaaccttgggttcca-3'-TAMRA	27	3645	364
Reverse	5'-tgactgttgtcttggcaatgt-3'	21	3681	365

10

Table AFB. Probe Name Ag5243

Primers	Sequence	Length	Start Position	SEQ ID No
Forward	5'-gactccttccaaaggctatattgt-3'	24	8809	366
Probe	TET-5'-cgattcaaggccctacaaatatctgcca-3'-TAMRA	28	8849	367
Reverse	5'-ccatttctggttccgtgtcta-3'	21	8880	368

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Table AFC.
Probe Name g5244

Primers	Sequences	Length	Start Position	SEQ ID No
Forward	5'-actgataattctattcctgaactgga-3'	26	4927	369
Probe	TET-5'-agctctgctagatctatctacagatataacgctgtaaaatc-3'-TAMRA	41	4992	370
Reverse	5'-aactcattatagatcatccaaaagga-3'	26	5036	371

20

Table AFD.
Probe Name g5245

Primers	Sequences	Length	Start Position	SEQ ID No

Forward	5'-accttgttgatgactttgctaatg-3'	24	4320	372
Probe	TET-5'-cagtggaactattacattccttccttgg caga-3'-TAMRA	32	4345	373
Reverse	5'-ggaagcgacacttcaatcaaa-3'	21	4387	374

Table AFE. Probe Name Ag5247

Primers	Sequenes	Length	Start Position	SEQ ID No
Forward	5'-acttacgttggacttaccatgg-3'	22	8183	375
Probe	TET-5'-caacttcatttcctcccagactaggtat gagg-3'-TAMRA	32	8211	376
Reverse	5'-tcatttcatttgaagtgagcaa-3'	22	8263	377

Table AFF. Probe Name Ag5248

Primers	Sequenes	Length	Start Position	SEQ ID No
Forward	5'-accttgttgatgactttgctaatg-3'	24	4320	378
Probe	TET-5'-cagtggaactattacattccttccttgg caga-3'-TAMRA	32	4345	379
Reverse	5'-caagaacatatatattcagaacctctgatc-3'	30	4377	380

Table AFG. AI_comprehensive panel_v1.0

Tissue Name	Rel. Exp.(%) Ag5242, Run 305464510	issue Name	Rel. Exp.(%) Ag5242, Run 305464510
110967 COPD-F	0.1	112427 Match Control Psoriasis-F	2.3
110980 COPD-F	1.1	112418 Psoriasis-M	0.1
110968 COPD-M	0.1	112723 Match Control Psoriasis-M	0.5
110977 COPD-M	4.4	112419 Psoriasis-M	0.0
110989 Emphysema-F	0.2	112424 Match Control Psoriasis-M	0.2
110992 Emphysema-F	2.7	112420 Psoriasis-M	1.8
110993 Emphysema-F	0.1	112425 Match Control Psoriasis-M	3.7
110994 Emphysema-F	0.1	104689 (MF) OA Bone-Backus	0.2
110995 Emphysema-F	6.8	104690 (MF) Adj "Normal" Bone-Backus	0.6

110996 Emphysema-F	2.0	104691 (MF) OA Synovium-Backus	0.1
110997 Asthma-M	0.1	104692 (BA) OA Cartilage-Backus	0.0
111001 Asthma-F	0.5	104694 (BA) OA Bone-Backus	0.2
111002 Asthma-F	0.9	104695 (BA) Adj "Normal" Bone-Backus	0.4
111003 Atopic Asthma-F	1.5	104696 (BA) OA Synovium-Backus	0.1
111004 Atopic Asthma-F	6.1	104700 (SS) OA Bone-Backus	0.9
111005 Atopic Asthma-F	2.5	104701 (SS) Adj "Normal" Bone-Backus	0.6
111006 Atopic Asthma-F	0.9	104702 (SS) OA Synovium-Backus	0.2
111417 Allergy-M	0.8	117093 OA Cartilage Rep7	0.9
112347 Allergy-M	0.0	112672 OA Bone5	0.0
112349 Normal Lung-F	0.0	112673 OA Synovium5	0.1
112357 Normal Lung-F	1.0	112674 OA Synovial Fluid cells5	0.2
112354 Normal Lung-M	0.7	117100 OA Cartilage Rep14	0.0
112374 Crohns-F	0.5	112756 OA Bone9	100.0
112389 Match Control Crohns-F	0.2	112757 OA Synovium9	6.4
112375 Crohns-F	0.1	112758 OA Synovial Fluid Cells9	0.1
112732 Match Control Crohns-F	0.3	117125 RA Cartilage Rep2	0.0
112725 Crohns-M	0.1	113492 Bone2 RA	31.6
112387 Match Control Crohns-M	0.1	113493 Synovium2 RA	11.8
112378 Crohns-M	0.0	113494 Syn Fluid Cells RA	22.2
112390 Match Control Crohns-M	1.5	113499 Cartilage4 RA	22.7
112726 Crohns-M	1.2	113500 Bone4 RA	28.1
112731 Match Control Crohns-M	0.9	113501 Synovium4 RA	20.2
112380 Ulcer Col-F	1.0	113502 Syn Fluid Cells4 RA	16.4
112734 Match Control Ulcer Col-F	0.8	113495 Cartilage3 RA	22.7
112384 Ulcer Col-F	3.7	113496 Bone3 RA	24.5
112737 Match Control Ulcer Col-F	0.8	113497 Synovium3 RA	14.7
112386 Ulcer Col-F	0.2	113498 Syn Fluid Cells3 RA	33.0
112738 Match Control Ulcer Col-F	0.5	117106 Normal Cartilage Rep20	0.0
112381 Ulcer Col-M	0.0	113663 Bone3 Normal	0.0
112735 Match Control Ulcer Col-M	0.0	113664 Synovium3 Normal	0.0
112382 Ulcer Col-M	0.3	113665 Syn Fluid Cells3 Normal	0.0
112394 Match Control Ulcer Col-M	0.1	117107 Normal Cartilage Rep22	0.1
112383 Ulcer Col-M	4.5	113667 Bone4 Normal	0.4
112736 Match Control Ulcer Col-M	0.3	113668 Synovium4 Normal	0.1

| 112423 Psoriasis-F | 0.2 | 113669 Syn Fluid Cells4 Normal | 0.8 |

Table AFH. CNS_neurodegeneration_v1.0

Tissue Name	Rel. Exp.(%) Ag5242, Run 229661546	Rel. Exp.(%) Ag5242, Run 233609876	Rel. Exp.(%) Ag5243, Run 229661547	Rel. Exp.(%) Ag5243, Run 276863566	Rel. Exp.(%) Ag5243, Run 277731460	Rel. Exp.(%) Ag5244, Run 229661548	Rel. Exp.(%) Ag5244, Run 233610762	Rel. Exp.(%) Ag5244, Run 277731461	Rel. Exp.(%) Ag5245, Run 229661549	Rel. Exp.(%) Ag5245, Run 230510320	Rel. Exp.(%) Ag5247, Run 229661550	Rel. Exp.(%) Ag5247, Run 276863570	Rel. Exp.(%) Ag5248, Run 229661551	Rel. Exp.(%) Ag5248, Run 276863572	Rel. Exp.(%) Ag5248, Run 277731466
AD 1 Hippo	22.4	21.6	29.3	31.6	27.5	9.1	0.0	3.1	16.0	0.0	9.0	6.7	14.9	13.7	17.9
AD 2 Hippo	47.3	42.0	54.7	53.2	44.8	0.0	2.9	4.0	16.2	4.6	41.8	21.8	44.4	32.8	32.5
AD 3 Hippo	12.2	13.5	17.8	13.6	10.9	0.0	0.0	0.0	0.0	0.0	5.8	0.0	9.8	4.8	6.8
AD 4 Hippo	14.8	14.4	16.6	17.7	20.6	0.0	0.0	0.0	23.2	7.6	17.3	8.6	12.8	6.4	7.0
AD 5 Hippo	65.5	84.1	61.6	63.7	57.4	6.7	0.0	4.3	11.6	5.3	84.7	31.0	85.3	61.1	62.0
AD 6 Hippo	56.3	59.5	82.4	84.7	90.1	74.2	57.8	51.8	58.6	30.8	100.0	92.0	69.3	48.3	55.5
Control 2 Hippo	29.5	25.7	29.3	31.6	31.9	0.0	0.0	5.5	15.1	29.1	42.0	29.9	27.7	25.0	26.1
Control 4 Hippo	32.8	29.7	35.6	31.2	37.1	8.1	11.3	0.0	0.0	0.0	27.0	23.7	25.2	20.9	16.5

Control (Path) 3 Hippo	33.9	33.9	24.7	24.0	30.8	0.0	4.5	0.0	8.2	0.0	13.0	12.2	13.1	19.8	22.1
AD 1 Temporal Ctx	32.3	33.9	32.3	34.6	35.4	2.0	5.4	2.9	9.3	19.5	29.9	21.9	26.2	17.9	26.1
AD 2 Temporal Ctx	35.8	42.3	39.5	51.1	46.3	3.3	5.4	0.0	14.2	0.0	28.7	32.5	38.2	37.6	**100.0**
AD 3 Temporal Ctx	28.3	21.2	20.4	23.5	20.7	0.0	0.0	0.0	0.0	0.0	4.4	4.5	12.2	9.5	11.3
AD 4 Temporal Ctx	47.3	44.8	36.6	39.0	45.4	10.3	0.0	8.3	39.0	19.2	43.5	25.3	33.0	25.9	29.5
AD 5 Inf Temporal Ctx	73.7	**100.0**	100.0	**100.0**	100.0	0.0	11.4	17.6	24.5	0.0	43.5	74.7	76.8	**100.0**	79.6
AD 5 Sup Temporal Ctx	93.3	77.4	87.7	82.4	88.3	7.3	10.7	8.9	29.1	3.3	45.7	59.0	82.4	70.7	64.2
AD 6 Inf Temporal Ctx	59.0	58.2	62.0	0.0	58.2	55.9	94.0	49.0	55.5	**100.0**	87.1	87.7	71.7	46.3	65.1

AD 6 Sup Temporal Ctx	85.3	99.3	74.2	74.7	90.1	100.0	100.0	100.0	99.3	73.7	95.9	97.3	97.3	60.3	94.0
Control 1 Temporal Ctx	47.6	46.3	27.4	28.5	29.1	1.7	0.0	0.0	58.2	27.7	25.3	19.1	44.4	25.2	32.3
Control 2 Temporal Ctx	37.6	37.4	30.6	27.5	32.8	2.7	11.0	4.5	31.4	48.3	8.9	15.4	50.0	34.4	29.1
Control 3 Temporal Ctx	27.5	24.1	27.4	32.8	37.6	7.1	5.4	2.6	5.1	6.3	16.6	5.8	35.6	21.3	27.7
Control 3 Temporal Ctx	38.2	39.0	34.6	30.6	31.9	8.7	2.6	4.9	8.4	0.0	31.4	13.7	22.4	26.1	31.4
Control (Path) 1 Temporal Ctx	66.0	81.2	54.0	58.6	52.5	2.5	0.0	3.2	78.5	37.9	72.7	72.7	75.8	63.3	69.7
Control (Path) 2 Temporal Ctx	43.5	50.0	40.1	41.8	41.5	2.0	10.9	0.0	80.1	20.9	42.6	31.9	42.9	33.9	42.0

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Control (Path) 3 Temporal Ctx	23.3	24.5	19.9	21.5	22.4	2.9	2.3	7.7	0.0	4.1	5.8	4.3	21.0	14.5	16.2
Control (Path) 4 Temporal Ctx	52.5	48.0	33.7	39.8	39.0	0.0	4.7	4.3	49.3	43.2	73.7	49.3	40.6	32.5	47.6
AD 1 Occipital Ctx	18.0	18.8	22.8	25.7	24.3	0.0	3.0	0.0	0.0	0.0	10.2	13.8	19.9	12.8	14.3
AD 2 Occipital Ctx (Missing)	0.0	0.0	0.0	0.0	0.0	0.0	0.0	0.0	0.0	0.0	0.0	0.0	0.0	0.0	0.0
AD 3 Occipital Ctx	15.5	14.0	17.8	17.8	18.0	0.0	0.0	0.0	3.2	0.0	10.2	0.0	10.3	5.2	5.5
AD 4 Occipital Ctx	17.3	23.7	25.3	27.5	24.3	3.3	3.1	3.3	28.7	6.7	22.2	23.0	8.6	17.0	21.5
AD 5 Occipital Ctx	22.4	26.1	21.3	15.2	22.5	2.0	3.1	5.3	25.7	5.1	16.7	8.7	3.3	20.9	18.4

AD 6 Occipital Ctx	28.9	21.6	19.1	20.4	18.9	11.7	15.6	2.8	0.0	18.2	12.1	0.0	29.9	18.0	24.7
Control 1 Occipital Ctx	9.3	10.2	6.8	6.1	7.4	0.0	0.0	0.0	4.3	7.8	5.6	0.0	4.8	3.7	3.3
Control 2 Occipital Ctx	34.9	33.0	24.7	28.7	31.6	0.0	5.1	7.4	31.2	17.9	7.9	4.6	39.5	20.2	28.3
Control 3 Occipital Ctx	27.2	24.1	27.5	25.2	24.5	2.4	9.2	4.2	7.0	0.0	13.8	0.0	14.5	17.6	16.8
Control 4 Occipital Ctx	19.6	20.3	18.0	26.8	21.2	0.0	1.6	0.0	0.0	0.0	12.5	5.6	15.4	8.8	12.9
Control (Path) 1 Occipital Ctx	56.6	64.6	48.6	58.6	57.8	9.1	5.1	7.8	66.4	30.6	69.7	57.0	76.3	42.6	55.5
Control (Path) 2 Occipital Ctx	5.7	6.1	9.0	7.1	8.5	2.0	0.0	0.0	5.6	0.0	1.6	0.0	16.3	3.8	4.1

Control (Path) 3 Occipital Ctx	2.6	3.1	4.1	1.9	4.5	0.0	0.0	0.0	0.0	0.0	0.0	0.0	1.5	1.8	1.7
Control (Path) 4 Occipital Ctx	9.9	11.2	5.4	9.2	8.2	0.0	0.0	0.0	11.7	15.7	5.1	7.2	2.1	0.3	5.0
Control 1 Parietal Ctx	28.9	32.5	19.6	21.8	22.4	0.0	0.0	0.0	0.0	3.6	9.8	16.4	23.8	16.3	17.1
Control 2 Parietal Ctx	100.0	90.8	79.0	83.5	76.3	7.9	23.8	9.7	26.8	12.2	39.0	37.9	100.0	44.1	63.3
Control 3 Parietal Ctx	14.8	11.9	17.3	15.3	17.0	0.0	9.8	0.0	0.0	0.0	1.7	7.2	12.9	8.5	11.3
Control (Path) 1 Parietal Ctx	62.4	68.3	57.8	70.2	63.7	4.2	3.8	0.0	100.0	55.9	41.5	100.0	99.3	53.2	71.2
Control (Path) 2 Parietal Ctx	17.1	19.8	22.1	21.0	25.9	1.9	10.4	0.0	30.8	0.0	17.9	18.0	6.3	10.2	15.8

Control (Path) 3 Parietal Ctx	12.0	10.2	11.7	8.4	13.9	2.8	5.3	0.0	6.3	0.0	3.9	0.0	3.2	4.9	4.2
Control (Path) 4 Parietal Ctx	30.1	25.5	26.1	25.7	29.1	1.5	0.0	0.0	59.0	40.9	23.7	30.1	25.9	26.4	17.8

Table AFI. General screening panel v1.5

5

Tissue Name	Rel. Exp.(%) Ag5242, Run 229665046	Rel. Exp.(%) Ag5243, Run 229665047	Rel. Exp (%) Ag5245, Run 229665049	Rel. Exp.(%) Ag5247, Run 229665052	Rel. Exp.(%) Ag5248, Run 229665053	Tissue Name	Rel. Exp.(%) Ag5242, Run 229665046	Rel. Exp.(%) Ag5243, Run 229665047	Rel. Exp.(%) Ag5245, Run 229665049	Rel. Exp.(%) Ag5247, Run 229665052	Rel. Exp.(%) Ag5248, Run 229665053
Adipose	0.1	0.0	0.0	0.0	0.0	Renal ca. TK-10	0.0	0.1	0.0	0.0	0.0
Melanoma* Hs688(A).T	0.8	0.5	0.0	0.0	1.2	Bladder	2.6	1.8	0.0	2.5	3.7
Melanoma* Hs688(B).T	0.1	0.0	0.0	0.0	0.0	Gastric ca. (liver met.) NCI-N87	0.0	0.0	0.0	0.0	0.0
Melanoma* M14	0.2	0.3	0.0	0.0	0.1	Gastric ca. KATO III	0.0	0.0	0.0	0.0	0.1
Melanoma* LOXIMVI	0.9	0.2	0.0	0.0	0.1	Colon ca. SW-948	5.2	4.6	0.4	0.6	3.7
Melanoma* SK-MEL-5	0.6	1.6	0.0	1.2	0.0	Colon ca. SW480	4.6	3.7	0.0	1.1	5.9
Squamous cell carcinoma SCC-4	0.0	0.0	0.0	0.0	0.0	Colon ca.* (SW480 met) SW620	0.1	0.0	0.0	0.0	0.0

Testis Pool	2.9	3.4	0.0	3.3	3.8	Colon ca. HT29	0.0	0.0	0.0	0.0	0.0
Prostate ca.* (bone met) PC-3	89.5	86.5	5.8	18.2	**100.0**	Colon ca. HCT-116	12.2	11.9	0.4	4.4	14.2
Prostate Pool	10.7	8.5	0.7	2.4	7.0	Colon ca. CaCo-2	13.8	14.0	8.9	6.6	16.4
Placenta	0.0	0.1	0.0	1.0	0.1	Colon cancer tissue	0.0	0.0	0.0	0.0	0.0
Uterus Pool	0.0	0.1	0.0	0.0	0.1	Colon ca. SW1116	0.1	0.0	0.0	0.0	0.0
Ovarian ca. OVCAR-3	10.5	18.7	12.1	1.7	16.7	Colon ca. Colo-205	0.0	0.0	0.0	0.0	1.2
Ovarian ca. SK-OV-3	0.2	0.1	0.0	0.2	0.0	Colon ca. SW-48	0.0	0.0	0.0	0.0	0.0
Ovarian ca. OVCAR-4	0.1	0.0	0.0	0.0	0.1	Colon Pool	0.1	0.0	0.0	0.6	0.1
Ovarian ca. OVCAR-5	7.3	7.1	0.0	3.7	12.1	Small Intestine Pool	3.7	1.6	1.6	1.0	4.1
Ovarian ca. IGROV-1	1.4	3.5	0.0	0.0	0.5	Stomach Pool	1.6	0.7	0.0	0.4	0.9
Ovarian ca. OVCAR-8	8.5	13.0	0.9	0.5	10.7	Bone Marrow Pool	0.1	0.0	0.0	0.0	0.1
Ovary	0.1	0.4	0.0	0.0	1.0	Fetal Heart	0.0	0.0	0.0	0.3	0.0
Breast ca. MCF-7	11.1	10.2	0.0	3.6	16.4	Heart Pool	0.1	0.0	0.0	0.7	0.1
Breast ca. MDA-MB-231	3.7	4.8	3.2	0.6	5.9	Lymph Node Pool	0.5	0.0	0.0	0.6	0.1
Breast ca. BT 549	0.0	0.0	0.0	0.0	0.0	Fetal Skeletal Muscle	0.2	0.0	0.0	1.1	0.0
Breast ca. T47D	10.2	4.4	0.0	3.1	9.9	Skeletal Muscle Pool	0.0	0.1	0.0	0.8	0.1
Breast ca. MDA-N	0.1	0.2	0.0	0.0	0.5	Spleen Pool	1.5	0.1	0.5	2.3	0.6

Breast Pool	0.8	1.9	0.0	0.9	1.5	Thymus Pool	3.2	1.7	1.9	0.7	2.9
Trachea	7.4	6.4	0.9	20.3	9.5	CNS cancer (glio/astro) U87-MG	4.4	2.6	0.3	1.2	3.2
Lung	0.3	0.0	0.0	0.0	0.1	CNS cancer (glio/astro) U-118-M G	0.1	0.0	0.0	0.0	0.0
Fetal Lung	25.7	20.9	1.7	6.7	22.2	CNS cancer (neuro;met) SK-N-AS	0.0	0.0	0.0	0.0	0.0
Lung ca. NCI-N417	3.4	3.6	0.0	0.7	11.6	CNS cancer (astro) SF-539	0.2	0.0	0.0	0.0	0.1
Lung ca. LX-1	0.1	0.0	0.0	0.0	0.0	CNS cancer (astro) SNB-75	0.1	0.1	0.0	0.0	0.2
Lung ca. NCI-H146	26.1	28.9	27.9	7.7	24.7	CNS cancer (glio) SNB-19	2.0	4.1	0.0	0.6	3.4
Lung ca. SHP-77	100.0	100.0	100.0	42.9	98.6	CNS cancer (glio) SF-295	2.4	3.3	0.4	0.3	4.1
Lung ca. A549	0.9	1.3	0.0	0.0	1.1	Brain (Amygdala) Pool	13.4	29.1	1.8	4.2	14.6
Lung ca. NCI-H526	1.8	1.1	0.0	0.0	1.9	Brain (cerebellum)	14.2	13.4	0.8	6.1	15.6
Lung ca. NCI-H23	0.0	0.0	0.0	0.0	0.2	Brain (fetal)	89.5	100.0	15.1	100.0	93.3
Lung ca. NCI-H460	5.4	3.3	9.3	48.3	23.5	Brain (Hippocampus) Pool	35.4	47.3	6.6	13.7	31.9
Lung ca. HOP-62	7.0	8.8	0.0	0.0	8.4	Cerebral Cortex Pool	40.1	53.2	8.9	35.1	39.0
Lung ca. NCI-H522	0.0	0.0	0.0	0.0	0.0	Brain (Substantia nigra) Pool	14.2	33.7	4.7	2.2	16.7

Liver	0.0	0.0	0.0	0.0	0.0	Brain (Thalamus) Pool	37.9	43.2	0.8	25.5	45.1
Fetal Liver	0.0	0.0	0.0	0.6	0.2	Brain (whole)	13.9	25.7	2.1	13.4	18.6
Liver ca. HepG2	0.0	0.1	0.0	0.0	0.0	Spinal Cord Pool	2.2	2.6	1.7	1.4	2.4
Kidney Pool	1.0	1.0	0.0	0.4	1.6	Adrenal Gland	0.7	0.7	0.8	1.9	0.3
Fetal Kidney	8.5	6.9	1.0	6.5	9.2	Pituitary gland Pool	18.6	16.3	5.0	3.2	36.6
Renal ca. 786-0	0.0	0.0	0.0	0.0	0.0	Salivary Gland	0.1	0.5	0.0	0.0	0.1
Renal ca. A498	0.1	0.0	0.0	0.0	0.0	Thyroid (female)	11.6	12.2	0.2	0.7	9.4
Renal ca. ACHN	0.4	0.1	0.0	0.0	0.5	Pancreatic ca. CAPAN2	0.1	0.0	0.0	0.0	0.1
Renal ca. UO-31	0.0	0.1	0.0	0.0	0.0	Pancreas Pool	3.6	2.3	0.0	1.7	2.0

Table AFJ. General screening panel v1.6

5

Tissue Name	Rel. Exp.(%) Ag5243, Run 277218719	Rel. Ex.(%) Ag5243, Run 277729929	Rel. Exp.(%) Ag5245, Run 277219697	Rel. Exp.(%) Ag5245, Run 277730879	Rel. Exp.(%) Ag5247, Run 277219699	Rel. Exp.(%) Ag5247, Run 277729933	Rel. Exp.(%) Ag5248, Run 277219701	Rel. Exp.(%) Ag5248, Run 277730881
Adipose	0.1	0.2	0.0	0.0	0.0	0.0	0.1	0.1
Melanoma* Hs688(A).T	0.0	0.0	0.0	0.0	0.0	0.0	0.0	0.0
Melanoma* Hs688(B).T	0.0	0.0	0.0	0.0	0.0	0.0	0.2	0.0
Melanoma* M14	0.2	0.0	0.7	0.0	0.0	0.0	0.0	0.3
Melanoma* LOXIMVI	0.2	0.1	0.0	0.0	0.0	0.0	0.1	0.2
Melanoma* SK-MEL-5	2.5	1.3	0.0	0.0	0.0	0.9	0.1	0.4
Squamous cell carcinoma SCC-4	0.0	0.0	0.0	0.0	0.0	0.0	0.0	0.1
Testis Pool	2.2	3.4	3.1	2.3	7.1	3.5	2.7	2.8
Prostate ca.* (bone met) PC-3	95.3	76.8	11.5	1.3	23.7	20.3	76.8	63.3

Prostate Pool	6.8	7.5	0.0	0.0	6.3	8.7	6.1	7.0
Placenta	0.0	0.0	0.0	0.0	0.0	0.0	0.1	0.1
Uterus Pool	0.0	0.0	0.0	0.0	0.0	0.0	0.0	0.1
Ovarian ca. OVCAR-3	13.2	11.7	9.5	4.0	3.3	5.2	11.6	14.5
Ovarian ca. SK-OV-3	0.2	0.3	0.0	0.0	0.0	0.0	0.1	0.3
Ovarian ca. OVCAR-4	0.0	0.0	0.0	0.0	0.0	0.0	0.0	0.0
Ovarian ca. OVCAR-5	6.6	7.4	2.3	0.0	4.7	0.8	4.7	5.1
Ovarian ca. IGROV-1	2.0	2.8	0.7	0.0	0.0	0.0	1.1	3.3
Ovarian ca. OVCAR-8	14.2	8.1	3.6	0.0	7.5	8.1	8.2	13.4
Ovary	0.1	0.6	0.0	0.0	0.0	0.0	0.7	0.2
Breast ca. MCF-7	7.4	8.0	0.0	0.0	3.5	9.4	8.0	9.2
Breast ca. MDA-MB-231	6.5	3.0	2.4	2.5	0.4	0.7	4.1	6.4
Breast ca. BT 549	0.0	0.0	0.0	0.0	0.0	1.0	0.0	0.0
Breast ca. T47D	6.7	3.8	0.8	0.0	5.5	1.5	4.7	8.0
Breast ca. MDA-N	0.0	0.2	0.5	0.0	0.0	0.5	0.1	0.3
Breast Pool	0.2	0.1	0.9	0.0	0.0	0.0	0.5	0.3
Trachea	18.6	15.6	3.9	0.0	14.6	18.0	5.5	7.6
Lung	0.2	0.0	0.0	0.0	0.0	1.2	0.0	0.1
Fetal Lung	21.3	21.0	0.0	0.7	10.3	5.1	19.3	23.7
Lung ca. NCI-N417	6.3	3.2	0.0	0.0	1.7	4.2	2.4	2.0
Lung ca. LX-1	0.0	0.0	0.0	0.0	0.0	0.0	0.0	0.0
Lung ca. NCI-H146	23.3	20.4	17.0	**100.0**	7.1	9.8	16.8	16.4
Lung ca. SHP-77	95.9	77.9	**100.0**	35.6	24.7	31.9	**100.0**	76.3
Lung ca. A549	1.0	0.4	0.0	0.0	0.0	0.0	0.3	1.1
Lung ca. NCI-H526	1.4	1.9	0.0	0.0	0.0	0.0	0.7	0.5
Lung ca. NCI-H23	0.0	0.0	0.0	0.0	0.0	0.0	0.0	0.1
Lung ca. NCI-H460	2.8	2.1	0.0	0.0	0.9	0.9	3.1	3.4
Lung ca. HOP-62	12.4	6.5	0.0	0.0	0.6	0.0	9.4	11.6

Lung ca. NCI-H522	0.0	0.0	0.0	0.0	0.0	0.0	0.0
Liver	0.0	0.0	0.0	0.0	0.0	0.0	0.0
Fetal Liver	0.0	0.0	0.0	0.0	0.0	0.9	0.2	0.0
Liver ca. HepG2	0.2	0.0	0.0	0.0	0.0	0.0	0.0	0.1
Kidney Pool	0.5	0.9	0.0	0.0	1.0	0.0	0.6	1.8
Fetal Kidney	5.8	6.8	0.0	0.0	11.4	6.6	4.3	7.9
Renal ca. 786-0	0.0	0.0	0.0	0.0	0.0	0.0	0.0	0.0
Renal ca. A498	0.0	0.0	0.0	0.0	0.0	0.0	0.0	0.0
Renal ca. ACHN	0.0	0.2	0.0	0.0	0.0	0.0	0.2	0.1
Renal ca. UO-31	0.2	0.2	0.0	0.0	0.0	0.0	0.0	0.1
Renal ca. TK-10	0.0	0.0	0.0	0.0	0.0	0.0	0.0	0.0
Bladder	1.2	1.5	0.0	0.0	3.8	1.4	3.3	3.2
Gastric ca. (liver met.) NCI-N87	0.0	0.0	0.0	0.0	0.0	0.0	0.0	0.0
Gastric ca. KATO III	0.0	0.0	0.0	0.0	0.0	0.0	0.0	0.0
Colon ca. SW-948	4.0	4.4	0.7	0.0	2.8	0.6	3.6	3.8
Colon ca. SW480	3.6	4.0	0.5	0.0	0.0	2.3	2.7	4.2
Colon ca.* (SW480 met) SW620	0.2	0.0	0.0	0.0	0.0	0.0	0.0	0.0
Colon ca. HT29	0.0	0.0	0.0	0.0	0.0	0.0	0.0	0.0
Colon ca. HCT-116	13.8	12.7	1.0	0.0	6.8	3.1	5.6	14.7
Colon ca. CaCo-2	18.8	14.9	10.8	4.7	10.2	10.1	2.4	11.6
Colon cancer tissue	0.0	0.0	0.0	0.0	0.0	0.0	0.0	0.1
Colon ca. SW1116	0.0	0.0	0.0	0.0	0.0	0.0	0.0	0.0
Colon ca. Colo-205	0.0	0.0	0.0	0.0	0.0	0.0	0.0	0.0
Colon ca. SW-48	0.0	0.0	0.0	0.0	0.0	0.0	0.0	0.0
Colon Pool	0.1	0.0	0.0	0.0	0.9	0.0	0.4	0.1
Small Intestine Pool	0.7	1.4	1.6	1.6	0.7	3.0	8.9	1.7

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Stomach Pool	0.6	1.0	0.0	0.0	0.0	0.0	0.7	0.6
Bone Marrow Pool	0.0	0.1	0.0	0.0	0.0	0.6	0.0	0.1
Fetal Heart	0.0	0.0	0.0	0.0	0.0	0.0	0.0	0.1
Heart Pool	0.0	0.0	0.0	0.0	0.0	0.0	0.0	0.0
Lymph Node Pool	0.0	0.7	0.0	0.0	0.8	0.0	0.5	0.4
Fetal Skeletal Muscle	0.4	0.1	0.0	0.0	0.0	0.0	0.1	0.2
Skeletal Muscle Pool	0.0	0.0	0.0	0.0	0.0	0.0	0.0	0.0
Spleen Pool	0.0	0.1	0.6	0.0	1.4	0.0	0.6	0.5
Thymus Pool	2.0	2.1	1.0	0.7	1.4	2.6	1.9	3.2
CNS cancer (glio/astro) U87-MG	2.6	2.5	0.8	0.0	0.7	0.6	3.7	4.3
CNS cancer (glio/astro) U-118-MG	0.3	0.1	0.0	0.0	0.0	0.0	0.0	0.0
CNS cancer (neuro;met) SK-N-AS	0.0	0.0	0.0	0.0	0.0	0.0	0.0	0.0
CNS cancer (astro) SF-539	0.0	0.0	0.0	0.0	0.0	0.0	0.1	0.1
CNS cancer (astro) SNB-75	0.2	0.3	0.0	0.0	0.0	0.0	0.5	0.3
CNS cancer (glio) SNB-19	3.1	2.4	0.0	0.0	0.0	1.1	1.9	3.4
CNS cancer (glio) SF-295	2.8	2.2	0.5	0.6	0.9	2.6	3.1	2.8
Brain (Amygdala) Pool	23.2	18.7	1.0	2.6	7.1	2.2	12.2	14.0
Brain (cerebellum)	13.8	11.7	3.1	1.0	10.2	11.3	13.3	14.1
Brain (fetal)	**100.0**	**100.0**	20.6	14.8	**100.0**	**100.0**	73.2	**100.0**
Brain (Hippocampus) Pool	51.1	40.3	6.9	5.3	25.9	14.3	26.8	35.8
Cerebral Cortex Pool	52.5	52.5	8.2	0.0	27.0	20.9	31.9	31.0
Brain (Substantia nigra) Pool	29.5	29.1	1.1	1.7	5.5	2.9	9.7	12.2
Brain (Thalamus) Pool	48.3	51.1	2.2	2.5	21.9	25.2	17.4	31.0
Brain (whole)	28.7	30.6	6.0	4.2	15.2	13.3	9.2	14.7

Spinal Cord Pool	1.9	1.3	1.3	0.0	1.0	0.0	1.6	2.2
Adrenal Gland	0.4	0.8	1.5	0.0	0.0	0.8	0.3	0.4
Pituitary gland Pool	17.9	13.7	2.6	7.4	0.0	11.1	13.4	15.8
Salivary Gland	0.2	0.3	0.0	0.0	0.0	0.0	0.3	0.6
Thyroid (female)	12.9	10.0	1.4	0.0	1.5	0.8	8.5	13.9
Pancreatic ca. CAPAN2	0.0	0.0	0.0	0.0	0.0	0.0	0.1	0.0
Pancreas Pool	2.6	3.2	0.0	0.0	0.6	3.6	4.5	3.7

Table AFK. Panel 4.1D

Tissue Name	Rel. Exp.() Ag5242, Run 229819771	Rel. Exp.(%) Ag5245, Run 229819577	Rel. Exp.(%) Ag5247, Run 229819792	Rel. Exp.(%) Ag5248, Run 229819793	Tissue Name	Rel. Exp.(%) Ag5242, Run 229819771	Rel. Exp.(%) Ag5245, Run 229819577	Rel. Exp.(%) Ag5247, Run 229819792	Rel. Exp.(%) Ag5248, Run 229819793
Secondary Th1 act	0.0	0.0	0.0	0.0	HUVEC IL-1beta	0.2	0.0	0.0	0.1
Secondary Th2 act	0.6	4.1	0.7	0.5	HUVEC IFN gamma	0.0	0.0	0.0	0.0
Secondary Tr1 act	2.3	1.2	0.6	2.3	HUVEC TNF alpha + IFN gamma	6.0	0.0	2.4	7.7
Secondary Th1 rest	0.0	0.0	0.0	0.1	HUVEC TNF alpha + IL4	1.0	0.0	0.6	4.2
Secondary Th2 rest	13.7	0.6	5.1	12.2	HUVEC IL-11	9.6	1.6	6.4	9.2
Secondary Tr1 rest	15.5	1.9	8.7	14.0	Lung Microvascular EC none	3.6	0.9	1.0	2.4
Primary Th1 act	100.0	71.7	100.0	85.3	Lung Microvascular EC TNFalpha + IL-1beta	0.0	0.0	0.0	0.0
Primary Th2 act	27.9	12.6	20.4	28.3	Microvascular Dermal EC none	0.1	0.0	0.0	0.3
Primary Tr1 act	36.6	9.4	24.3	28.9	Microsvasular Dermal EC TNFalpha + IL-1beta	0.0	0.0	0.0	0.0
Primary Th1 rest	15.9	2.9	5.1	14.6	Bronchial epithelium TNFalpha + IL1beta	0.1	0.0	0.0	0.0
Primary Th2 rest	34.2	3.4	23.3	29.1	Small airway epithelium none	0.2	0.0	0.0	0.2

Primary Tr1 rest	12.0	5.0	12.7	12.9	Small airway epithelium TNFalpha + IL-1beta	3.1	0.0	0.7	3.7
CD45RA CD4 lymphocyte act	0.6	0.0	0.0	0.0	Coronery artery SMC rest	4.1	0.0	0.6	3.6
CD45RO CD4 lymphocyte act	0.0	0.0	0.0	0.2	Coronery artery SMC TNFalpha + IL-1beta	3.1	0.0	0.0	2.6
CD8 lymphocyte act	5.6	2.9	0.7	7.3	Astrocytes rest	3.8	0.9	0.6	4.0
Secondary CD8 lymphocyte rest	0.0	0.0	0.0	0.0	Astrocytes TNFalpha + IL-1beta	0.0	0.0	0.0	0.0
Secondary CD8 lymphocyte act	2.1	0.0	0.0	1.9	KU-812 (Basophil) rest	0.0	0.0	0.0	0.0
CD4 lymphocyte none	8.1	1.2	5.8	7.4	KU-812 (Basophil) PMA/ionomycin	12.6	1.0	4.5	15.4
2ry Th1/Th2/Tr1_anti -CD95 CH11	0.0	0.0	0.0	0.0	CCD1106 (Keratinocytes) none	15.7	15.5	4.3	15.8
LAK cells rest	0.1	0.0	0.6	0.1	CCD1106 (Keratinocytes) TNFalpha + IL-1beta	0.0	0.0	0.0	0.0
LAK cells IL-2	0.3	0.0	0.0	0.3	Liver cirrhosis	0.1	0.0	0.0	0.0
LAK cells IL-2+IL-12	25.2	3.1	4.8	24.0	NCI-H292 none	0.0	0.0	0.0	0.0
LAK cells IL-2+IFN gamma	0.2	0.0	0.0	1.1	NCI-H292 IL-4	0.0	0.0	0.0	0.0
LAK cells IL-2+ IL-18	0.5	0.0	0.7	0.7	NCI-H292 IL-9	0.0	0.0	0.0	0.0
LAK cells PMA/ionomycin	0.2	0.0	0.6	0.0	NCI-H292 IL-13	0.2	0.0	0.0	0.0
NK Cells IL-2 rest	0.5	1.9	0.0	0.5	NCI-H292 IFN gamma	0.0	0.0	0.0	0.0
Two Way MLR 3 day	4.5	5.1	0.7	2.3	HPAEC none	0.0	0.0	0.0	0.0
Two Way MLR 5 day	6.7	14.9	9.5	15.0	HPAEC TNF alpha + IL-1 beta	0.1	0.0	0.0	0.0
Two Way MLR 7 day	0.2	0.0	0.0	0.1	Lung fibroblast none	0.0	0.0	0.0	0.0
PBMC rest	8.7	0.0	2.3	6.0	Lung fibroblast TNF alpha + IL-1 beta	19.9	25.7	4.4	22.8
PBMC PWM	0.2	0.0	0.0	0.4	Lung fibroblast IL-4	72.2	100.0	32.8	49.7
PBMC PHA-L	0.2	0.0	0.0	0.1	Lung fibroblast IL-9	1.2	0.0	0.4	0.6
Ramos (B cell) none	3.6	2.2	1.1	1.9	Lung fibroblast IL-13	1.8	0.0	1.5	1.2
Ramos (B cell) ionomycin	1.8	3.6	1.5	2.2	Lung fibroblast IFN gamma	0.0	0.0	0.0	0.0
B lymphocytes PWM	1.3	0.0	2.0	1.1	Dermal fibroblast CCD1070 rest	0.1	0.0	0.0	0.0

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B lymphocytes CD40L and IL-4	0.8	0.7	1.2	1.5	Dermal fibroblast CCD1070 TNF alpha	2.9	0.0	1.3	5.3
EOL-1 dbcAMP	3.7	6.7	3.3	2.0	Dermal fibroblast CCD1070 IL-1 beta	6.3	0.0	1.7	7.7
EOL-1 dbcAMP PMA/ionomycin	3.0	0.0	2.3	2.0	Dermal fibroblast IFN gamma	0.0	0.0	0.0	0.0
Dendritic cells none	10.7	1.9	3.8	13.6	Dermal fibroblast IL-4	0.0	0.0	0.0	0.0
Dendritic cells LPS	4.7	6.2	11.7	8.2	Dermal Fibroblasts rest	0.0	0.0	0.0	0.0
Dendritic cells anti-CD40	0.1	0.0	0.0	0.0	Neutrophils TNFa+LPS	0.1	0.0	0.0	0.0
Monocytes rest	11.6	0.6	2.8	16.4	Neutrophils rest	87.7	11.7	28.3	100.0
Monocytes LPS	4.6	5.6	1.4	5.4	Colon	0.0	0.0	0.0	0.0
Macrophages rest	0.2	0.0	0.0	0.1	Lung	0.2	0.0	0.0	0.3
Macrophages LPS	11.5	0.0	0.9	9.2	Thymus	0.1	0.0	0.0	0.6
HUVEC none	0.3	0.0	0.0	0.5	Kidney	0.1	0.0	1.4	0.6
HUVEC starved	15.9	8.4	2.4	15.5					

Table AFL. general oncology screening panel_v_2.4

Tissue Name	Rel. Exp.(%) Ag5242, Run 260269083	Rel. Exp.(%) Ag5247, Run 260269132	Rel. Exp.(%) Ag5248, Run 260269133	issue Name	Rel. Exp.(%) Ag5242, Run 260269083	Rel. Exp.(%) Ag5247, Run 260269132	Rel. Exp.(%) Ag5248, Run 260269133
Colon cancer 1	0.0	0.0	3.5	Bladder cancer NAT 2	0.0	0.0	0.0
Colon cancer NAT 1	7.2	0.0	11.0	Bladder cancer NAT 3	0.0	0.0	0.0
Colon cancer 2	0.0	0.0	0.0	Bladder cancer NAT 4	0.0	0.0	0.0
Colon cancer NAT 2	17.6	16.6	15.7	Prostate adenocarcinoma 1	2.4	20.9	5.8
Colon cancer 3	4.5	0.0	3.8	Prostate adenocarcinoma 2	0.0	0.0	2.0
Colon cancer NAT 3	37.1	0.0	27.0	Prostate adenocarcinoma 3	71.7	55.9	54.3
Colon malignant cancer 4	6.1	0.0	1.0	Prostate adenocarcinoma 4	1.0	0.0	7.2

Colon normal adjacent tissue 4	0.0	0.0	2.4	Prostate cancer NAT 5	4.5	0.0	0.0
Lung cancer 1	25.0	17.9	4.2	Prostate adenocarcinoma 6	30.6	4.5	11.1
Lung NAT 1	2.3	3.9	12.9	Prostate adenocarcinoma 7	14.4	6.3	23.0
Lung cancer 2	40.1	**100.0**	**100.0**	Prostate adenocarcinoma 8	9.1	5.0	6.8
Lung NAT 2	32.3	18.2	48.6	Prostate adenocarcinoma 9	75.3	10.7	31.0
Squamous cell carcinoma 3	73.2	47.0	82.4	Prostate cancer NAT 10	0.0	0.0	7.1
Lung NAT 3	13.3	3.5	5.8	Kidney cancer 1	0.0	0.0	0.0
metastatic melanoma 1	4.4	0.0	1.5	KidneyNAT 1	33.7	11.7	10.7
Melanoma 2	0.0	0.0	1.4	Kidney cancer 2	10.7	7.4	2.8
Melanoma 3	9.8	0.0	4.2	Kidney NAT 2	**100.0**	42.9	51.4
metastatic melanoma 4	2.1	0.0	1.0	Kidney cancer 3	61.1	8.6	24.8
metastatic melanoma 5	6.4	9.3	2.2	Kidney NAT 3	63.3	16.0	29.9
Bladder cancer 1	0.0	0.0	0.0	Kidney cancer 4	8.8	0.0	1.9
Bladder cancer NAT 1	0.0	0.0	0.0	Kidney NAT 4	5.3	0.0	9.2
Bladder cancer 2	2.1	0.0	0.0				

AI_comprehensive panel_v1.0 Summary: Ag5242 Highest expression is seen in osteoarthritic bone sample (CT=27.5). Prominenet levels of expression are seen in a cluster of samples derived from RA. Thus, expression of this gene could be used to differentiate between these samples and other samples on this panel and as a marker of rheumatoid arthritis. In addition, modulation of the expression or function of this gene may be useful in the treatment of RA.

CNS_neurodegeneration_v1.0 Summary: Ag5242/Ag5243/Ag5247/Ag5248 Multiple experiments with four different probe and primer sets produce results that are in reasonable agreement. These panels do not show differential expression of this gene in Alzheimer's disease. However, these profiles confirm the expression of this gene at

moderate levels in the brain. Please see Panel 1.5 for discussion of this gene in the central nervous system.

Ag5244 Three experiments with Ag5244, which is specific for CG150799-03, detect expression of this gene at low but significant levels in the hippocampus and temporal

5   cortex of Alzheimer's patients. This expression may suggest an involvement of this gene product in the etiology of this disease.

One experiment with Ag5244 (Run 276863567) and two experiments with Ag5245 (Run 276863569 and Run 277731463), also specific for CG150799-03, show low/undetectable levels of expression (CTs>35). (Data not shown). Two additional

10 experiments with Ag5245 show low expression in samples from the parietal cortex of a normal patient and the inferior temporal cortex of an Alzheimer's patient.

General_screening_panel_v1.5

Summary: Ag5242/Ag5243/Ag5245/Ag5247/Ag5248 Multiple experiments with five different probe and primer sets produce results that are in reasonable agreement. Highest

15 expression is seen in cell lines from lung and prostate cancers and the fetal brain (CTs=28-30). This gene, which encodes a MASS1 homolog, appears be preferentially expressed in the brain, with prominent levels of expression in all regions of the CNS examined. MASS1 is a large, calcium-binding GPCR expressed in the central nervous system that may play a fundamental role in its development (MacMillan, J Biol Chem 2002

20 Jan 4;277(1):785-92). In addition, this gene has been associated with some nonsymptomatic epilepsies (Skardski, Neuron, Vol 31, 537-544, August 2001). Thus, based on the homology of this protein to MASS1 and the preferential expression in the brain, expression of this gene could be used to differentiate between brain and non-neural tissue. In addition, therapeutic modulation of the expression or function of this gene may be useful

25 in the treatment of neurological disorders, such as Alzheimer's disease, Parkinson's disease, schizophrenia, multiple sclerosis, stroke and epilepsy.

Moderate levels of expression are also seen in samples from lung, colon, ovarian and prostate cancer cell lines. This suggests that expression of this gene could be used as a marker of these cancers. Futhermore, therapeutic modulation of the expression or function

30 of this gene may be useful in the treatment of these cancers.

Ag5244 Expression of this gene is low/undetectable (CTs > 35) across all of the samples on this panel.

General_screening_panel_v1.6 Summary: Ag5243/Ag5247/Ag5248/Ag5245
Multiple experiments with three different probe and primer sets produce results that are in very good agreement. Highest expression is seen in a lung cancer cell line and the fetal brain (CTs=27-32). Overall, expression is in excellent agreement with Panel 1.5, with
5   prominent expression seen in all regions of the CNS, and lung and prostate cancer cell lines. Please see Panel 1.5 for further discussion of this gene.

Ag5244 Expression of this gene is low/undetectable (CTs > 35) across all of the samples on this panel.

Panel 4.1D Summary: Ag5242/Ag5243/Ag5247/Ag5248 Multiple experiments
10   with four different probe and primers sets show highest expression of this gene in primary activated Th1 cells and resting neutrophils (CTs=27-31). Since this gene is expressed predominantly in activated Th-1 vs Th-2 cells, regulation of the expression of this gene might also be important for autoimmune disease such as rheumatoid arthritis (please see also AI panel). Moderate levels of expression are also seen in IL-4 treated lung fibroblasts
15   and resting neutrophils. Thus, therapeutic regulation of the transcript or the protein encoded by the transcript could be important in immune modulation and in the treatment of T cell-mediated diseases such as asthma, arthritis, psoriasis, IBD, and lupus.

Ag5245 Highest expression of this gene is seen in IL-4 treated lung fibroblasts (CT=32). Low but significant expression is also seen in TNF-a/IL1-b treated lung
20   fibroblasts and primary activated Th1 cells. Three experiments with the probe and primer set Ag5244 show low/undetectable levels of results (CTs>35).

general oncology screening panel_v_2.4
Summary: Ag5242/Ag5243/Ag5247/Ag5248 Four experiments with the different probe and primer sets show highest expression in a lung cancers and normal kidney tissue
25   adjacent to a tumor (CTs=31-34). Overall, this gene is expressed at low but significant levels in prostate cancer, normal kidney and kidney cancer, squamous cell carcinoma and normal colon. Therefore, therapeutic modulation of this gene or its protein product may be useful in the treatment of lung, prostate and kidney cancers.

Ag5244/Ag5245 Expression of this gene is low/undetectable in all samples on this
30   panel (CTs>35).

AG. CG151014-01: Metabotropic glutamate receptor 3-variant

Expression of gene CG151014-01 was assessed using the primer-probe set Ag5219, described in Table AGA. Results of the RTQ-PCR runs are shown in Tables AGB, AGC and AGD.

Table AGA. Probe Name Ag5219

Primers		Length	Start Position	SEQ ID No
Forward	5'-tgattgtgaattgcagttcagt-3'	22	2550	381
Probe	TET-5'-aagtgctcacgtgcagctccagaata-3'-TAMRA	26	2598	382
Reverse	5'-gtactagggttgttcttttgctct-3'	24	2631	383

Table AGB. CNS_neurodegeneration_v1.0

Tissue Name	Rel. Exp.(%) Ag5219, Run 228020421	issue Name	Rel. Exp.(%) Ag5219, Run 228020421
AD 1 Hippo	9.4	Control (Path) 3 Temporal Ctx	6.5
AD 2 Hippo	24.8	Control (Path) 4 Temporal Ctx	25.0
AD 3 Hippo	6.3	AD 1 Occipital Ctx	15.7
AD 4 Hippo	7.6	AD 2 Occipital Ctx (Missing)	0.0
AD 5 Hippo	53.2	AD 3 Occipital Ctx	6.8
AD 6 Hippo	24.1	AD 4 Occipital Ctx	33.2
Control 2 Hippo	40.9	AD 5 Occipital Ctx	51.8
Control 4 Hippo	6.7	AD 6 Occipital Ctx	15.3
Control (Path) 3 Hippo	5.6	Control 1 Occipital Ctx	7.6
AD 1 Temporal Ctx	19.1	Control 2 Occipital Ctx	46.0
AD 2 Temporal Ctx	34.9	Control 3 Occipital Ctx	16.6
AD 3 Temporal Ctx	5.6	Control 4 Occipital Ctx	8.5
AD 4 Temporal Ctx	25.3	Control (Path) 1 Occipital Ctx	90.1
AD 5 Inf Temporal Ctx	100.0	Control (Path) 2 Occipital Ctx	11.5
AD 5 Sup Temporal Ctx	32.5	Control (Path) 3 Occipital Ctx	3.8
AD 6 Inf Temporal Ctx	44.1	Control (Path) 4 Occipital Ctx	11.9
AD 6 Sup Temporal Ctx	32.5	Control 1 Parietal Ctx	9.5
Control 1 Temporal Ctx	10.5	Control 2 Parietal Ctx	40.6
Control 2 Temporal Ctx	45.4	Control 3 Parietal Ctx	18.3
Control 3 Temporal Ctx	28.9	Control (Path) 1 Parietal Ctx	74.2
Control 3 Temporal Ctx	10.1	Control (Path) 2 Parietal Ctx	27.5
Control (Path) 1 Temporal Ctx	65.1	Control (Path) 3 Parietal Ctx	5.0
Control (Path) 2 Temporal Ctx	36.1	Control (Path) 4 Parietal Ctx	36.3

Table AGC. General screening panel v1.5

5

Tissue Name	Rel. Exp.(%) Ag5219, Run 228758224	issue Name	Rel. Exp.(%) Ag5219, Run 228758224
Adipose	0.3	Renal ca. TK-10	0.4
Melanoma* Hs688(A).T	0.0	Bladder	0.2
Melanoma* Hs688(B).T	0.0	Gastric ca. (liver met.) NCI-N87	6.6
Melanoma* M14	0.0	Gastric ca. KATO III	0.0
Melanoma* LOXIMVI	0.5	Colon ca. SW-948	0.1
Melanoma* SK-MEL-5	0.8	Colon ca. SW480	0.6
Squamous cell carcinoma SCC-4	0.8	Colon ca.* (SW480 met) SW620	1.1
Testis Pool	0.4	Colon ca. HT29	0.0
Prostate ca.* (bone met) PC-3	2.1	Colon ca. HCT-116	1.7
Prostate Pool	0.5	Colon ca. CaCo-2	0.7
Placenta	0.0	Colon cancer tissue	0.0
Uterus Pool	0.2	Colon ca. SW1116	0.0
Ovarian ca. OVCAR-3	1.0	Colon ca. Colo-205	0.0
Ovarian ca. SK-OV-3	0.9	Colon ca. SW-48	0.0
Ovarian ca. OVCAR-4	0.0	Colon Pool	0.7
Ovarian ca. OVCAR-5	0.2	Small Intestine Pool	0.7
Ovarian ca. IGROV-1	0.0	Stomach Pool	1.4
Ovarian ca. OVCAR-8	0.1	Bone Marrow Pool	0.1
Ovary	0.1	Fetal Heart	0.6
Breast ca. MCF-7	0.0	Heart Pool	0.3
Breast ca. MDA-MB-231	0.5	Lymph Node Pool	1.1
Breast ca. BT 549	0.0	Fetal Skeletal Muscle	0.1
Breast ca. T47D	0.0	Skeletal Muscle Pool	0.7
Breast ca. MDA-N	0.0	Spleen Pool	1.4
Breast Pool	2.6	Thymus Pool	0.4
Trachea	0.4	CNS cancer (glio/astro) U87-MG	1.0
Lung	0.2	CNS cancer (glio/astro) U-118-MG	0.1
Fetal Lung	0.8	CNS cancer (neuro;met) SK-N-AS	1.4
Lung ca. NCI-N417	0.1	CNS cancer (astro) SF-539	0.0
Lung ca. LX-1	4.5	CNS cancer (astro) SNB-75	0.0
Lung ca. NCI-H146	1.1	CNS cancer (glio) SNB-19	0.0
Lung ca. SHP-77	3.3	CNS cancer (glio) SF-295	0.0
Lung ca. A549	0.0	Brain (Amygdala) Pool	60.3
Lung ca. NCI-H526	0.3	Brain (cerebellum)	**100.0**
Lung ca. NCI-H23	0.4	Brain (fetal)	66.4

Lung ca. NCI-H460	0.9	Brain (Hippocampus) Pool	43.5
Lung ca. HOP-62	0.0	Cerebral Cortex Pool	80.1
Lung ca. NCI-H522	0.7	Brain (Substantia nigra) Pool	54.0
Liver	0.0	Brain (Thalamus) Pool	94.6
Fetal Liver	0.4	Brain (whole)	65.1
Liver ca. HepG2	0.9	Spinal Cord Pool	36.6
Kidney Pool	1.5	Adrenal Gland	0.6
Fetal Kidney	0.7	Pituitary gland Pool	0.9
Renal ca. 786-0	0.0	Salivary Gland	0.2
Renal ca. A498	0.0	Thyroid (female)	0.0
Renal ca. ACHN	1.0	Pancreatic ca. CAPAN2	0.1
Renal ca. UO-31	0.5	Pancreas Pool	0.9

Table AGD. Panel 4.1D

5

Tissue Name	Rel. Exp (%) Ag5219, Run 229739298	Tissue Name	Rel. Exp.(%) Ag5219, Run 229739298
Secondary Th1 act	0.0	HUVEC IL-1beta	3.3
Secondary Th2 act	3.2	HUVEC IFN gamma	14.0
Secondary Tr1 act	0.0	HUVEC TNF alpha + IFN gamma	2.9
Secondary Th1 rest	0.0	HUVEC TNF alpha + IL4	1.8
Secondary Th2 rest	0.0	HUVEC IL-11	21.8
Secondary Tr1 rest	2.9	Lung Microvascular EC none	100.0
Primary Th1 act	0.0	Lung Microvascular EC TNFalpha + IL-1beta	31.9
Primary Th2 act	5.8	Microvascular Dermal EC none	0.0
Primary Tr1 act	0.0	Microsvasular Dermal EC TNFalpha + IL-1beta	15.5
Primary Th1 rest	0.0	Bronchial epithelium TNFalpha + IL1beta	0.0
Primary Th2 rest	1.8	Small airway epithelium none	0.0
Primary Tr1 rest	4.7	Small airway epithelium TNFalpha + IL-1beta	3.4
CD45RA CD4 lymphocyte act	0.0	Coronery artery SMC rest	2.3
CD45RO CD4 lymphocyte act	11.1	Coronery artery SMC TNFalpha + IL-1beta	0.0
CD8 lymphocyte act	6.7	Astrocytes rest	0.0
Secondary CD8 lymphocyte rest	5.9	Astrocytes TNFalpha + IL-1beta	3.4
Secondary CD8 lymphocyte act	0.0	KU-812 (Basophil) rest	4.1
CD4 lymphocyte none	3.3	KU-812 (Basophil) PMA/ionomycin	26.1

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2ry Th1/Th2/Tr1_anti-CD95 CH11	5.9	CCD1106 (Keratinocytes) none	4.5
LAK cells rest	3.0	CCD1106 (Keratinocytes) TNFalpha + IL-1beta	0.0
LAK cells IL-2	2.0	Liver cirrhosis	0.0
LAK cells IL-2+IL-12	0.0	NCI-H292 none	18.2
LAK cells IL-2+IFN gamma	3.0	NCI-H292 IL-4	16.7
LAK cells IL-2+ IL-18	2.7	NCI-H292 IL-9	25.0
LAK cells PMA/ionomycin	0.0	NCI-H292 IL-13	48.3
NK Cells IL-2 rest	24.1	NCI-H292 IFN gamma	19.9
Two Way MLR 3 day	3.5	HPAEC none	8.1
Two Way MLR 5 day	1.5	HPAEC TNF alpha + IL-1 beta	7.8
Two Way MLR 7 day	0.0	Lung fibroblast none	0.0
PBMC rest	0.0	Lung fibroblast TNF alpha + IL-1 beta	2.0
PBMC PWM	1.0	Lung fibroblast IL-4	7.9
PBMC PHA-L	0.0	Lung fibroblast IL-9	0.0
Ramos (B cell) none	18.2	Lung fibroblast IL-13	0.0
Ramos (B cell) ionomycin	59.9	Lung fibroblast IFN gamma	2.8
B lymphocytes PWM	4.2	Dermal fibroblast CCD1070 rest	0.0
B lymphocytes CD40L and IL-4	13.2	Dermal fibroblast CCD1070 TNF alpha	0.0
EOL-1 dbcAMP	0.0	Dermal fibroblast CCD1070 IL-1 beta	6.7
EOL-1 dbcAMP PMA/ionomycin	4.8	Dermal fibroblast IFN gamma	40.6
Dendritic cells none	4.4	Dermal fibroblast IL-4	25.0
Dendritic cells LPS	0.0	Dermal Fibroblasts rest	2.1
Dendritic cells anti-CD40	0.0	Neutrophils TNFa+LPS	0.0
Monocytes rest	0.0	Neutrophils rest	0.0
Monocytes LPS	0.0	Colon	0.0
Macrophages rest	0.0	Lung	0.0
Macrophages LPS	0.0	Thymus	0.0
HUVEC none	1.7	Kidney	11.3
HUVEC starved	28.1		

CNS_neurodegeneration_v1.0 Summary: Ag5219 This panel confirms the expression of this gene at low levels in the brain in an independent group of individuals. This gene is found to be slightly down-regulated in the temporal cortex of Alzheimer's disease patients. Therefore, up-regulation of this gene or its protein product, or treatment with specific agonists for this receptor may be of use in reversing the dementia, memory loss, and neuronal death associated with this disease.

General_screening_panel_v1.5 Summary: Ag5219 Highest expression of this

gene is deted in cerebellum (CT=27). High expression of this gene is mainly seen in all the

region of central nervous system examined, including amygdala, hippocampus, substantia

nigra, thalamus, cerebellum, cerebral cortex, and spinal cord. Therefore, therapeutic

5   modulation of this gene product may be useful in the treatment of central nervous system

disorders such as Alzheimer's disease, Parkinson's disease, epilepsy, multiple sclerosis,

schizophrenia and depression.

In·addition, moderate to low levels of expression of this gene is also seen in a

number of cancer cell lines derived from brain, colon, gastric, lung, ovarian, and prostate

10   cancers, squamous cell carcinoma and melanoma. Therefore, therapeutic modulation of this

gene may be useful in the treatment of these cancers.

Low levels of expression of this gene is also seen in tissues with

metabolic/endocrine functions including pancreas, adrenal and pituitary cancers, fetal heart,

skeletal muscle and gastrointestinal tract. Therefore, therapeutic modulation of the activity

15   of this gene may prove useful in the treatment of endocrine/metabolically related diseases,

such as obesity and diabetes.

Panel 4.1D Summary: Ag5219 Highest expression of this gene is detected in lung

microvascular endothelial cells (CT=32.4). This gene is expressed at lower levels in

cytokine activated lung microvascular cells, activated dermal fibroblasts, resting and

20   activated mucoepidermoid NCI-H292, activated basophils, starved and IL-11 stimulated

HUVEC cells, Ramos B cells, and resting IL-2 treated NK cells. Therefore, therapeutic

modulation of this gene may be useful in the treatment of autoimmune and inflammatory

diseases such as asthma, allergies, inflammatory bowel disease, lupus erythematosus,

psoriasis, rheumatoid arthritis, and osteoarthritis.

25   **AH. CG151014-02 and CG151014-03: Metabotropic glutamate**
     receptor 3.

Expression of gene CG151014-02 and CG151014-02 was assessed using the

primer-probe set Ag5220, described in Table AHA. Results of the RTQ-PCR runs are

shown in Tables AHB and AHC. Please note that CG151014-03 represents a full-length

30   physical clone.

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Table AHA. Probe Name Ag5220

Primers		Length	Start Position	SEQ ID No
Forward	5'-atcaacttcacgggtgcag-3'	19	1399	384
Probe	TET-5'-ctttgtggtcttgggctgtttgtttg-3'-TAMRA	26	1453	385
Reverse	5'-caggatgatgtgaaccttgg-3'	20	1482	386

5

Table AHB. CNS_neurodegeneration_v1.0

Tissue Name	Rel. Exp.(%) Ag5220, Run 228020422	issue Name	Rel. Exp.(%) Ag5220, Run 228020422
AD 1 Hippo	2.0	Control (Path) 3 Temporal Ctx	5.8
AD 2 Hippo	49.0	Control (Path) 4 Temporal Ctx	25.2
AD 3 Hippo	1.0	AD 1 Occipital Ctx	5.6
AD 4 Hippo	13.5	AD 2 Occipital Ctx (Missing)	0.0
AD 5 hippo	35.4	AD 3 Occipital Ctx	3.1
AD 6 Hippo	59.9	AD 4 Occipital Ctx	24.7
Control 2 Hippo	34.2	AD 5 Occipital Ctx	17.2
Control 4 Hippo	7.0	AD 6 Occipital Ctx	61.6
Control (Path) 3 Hippo	4.4	Control 1 Occipital Ctx	2.6
AD 1 Temporal Ctx	6.0	Control 2 Occipital Ctx	43.2
AD 2 Temporal Ctx	39.2	Control 3 Occipital Ctx	10.2
AD 3 Temporal Ctx	2.4	Control 4 Occipital Ctx	9.0
AD 4 Temporal Ctx	29.9	Control (Path) 1 Occipital Ctx	100.0
AD 5 Inf Temporal Ctx	76.3	Control (Path) 2 Occipital Ctx	7.7
AD 5 SupTemporal Ctx	29.9	Control (Path) 3 Occipital Ctx	2.1
AD 6 Inf Temporal Ctx	60.3	Control (Path) 4 Occipital Ctx	14.2
AD 6 Sup Temporal Ctx	69.3	Control 1 Parietal Ctx	7.0
Control 1 Temporal Ctx	13.2	Control 2 Parietal Ctx	24.3
Control 2 Temporal Ctx	52.9	Control 3 Parietal Ctx	15.4
Control 3 Temporal Ctx	23.3	Control (Path) 1 Parietal Ctx	89.5
Control 4 Temporal Ctx	11.7	Control (Path) 2 Parietal Ctx	15.2
Control (Path) 1 Temporal Ctx	87.1	Control (Path) 3 Parietal Ctx	6.4
Control (Path) 2 Temporal Ctx	59.0	Control (Path) 4 Parietal Ctx	33.0

10

Table AHC. General screening panel v1.5

Tissue Name	Rel. Exp.(%) Ag5220, Run 228758228	Tissue Name	Rel. Exp.(%) Ag5220, Run 228758228
Adipose	0.0	Renal ca. TK-10	0.0
Melanoma* Hs688(A).T	0.0	Bladder	0.0
Melanoma* Hs688(B).T	0.0	Gastric ca. (liver met.) NCI-N87	0.0
Melanoma* M14	0.0	Gastric ca. KATO III	0.0
Melanoma* LOXIMVI	0.0	Colon ca. SW-948	0.0
Melanoma* SK-MEL-5	0.0	Colon ca. SW480	0.0
Squamous cell carcinoma SCC-4	0.0	Colon ca.* (SW480 met) SW620	0.0
Testis Pool	0.0	Colon ca. HT29	0.0
Prostate ca.* (bone met) PC-3	0.0	Colon ca. HCT-116	0.0
Prostate Pool	0.0	Colon ca. CaCo-2	0.0
Placenta	0.0	Colon cancer tissue	0.0
Uterus Pool	0.0	Colon ca. SW1116	0.0
Ovarian ca. OVCAR-3	0.0	Colon ca. Colo-205	0.0
Ovarian ca. SK-OV-3	0.0	Colon ca. SW-48	0.0
Ovarian ca. OVCAR-4	0.0	Colon Pool	0.0
Ovarian ca. OVCAR-5	0.0	Small Intestine Pool	0.0
Ovarian ca. IGROV-1	0.0	Stomach Pool	1.6
Ovarian ca. OVCAR-8	0.0	Bone Marrow Pool	0.0
Ovary	0.0	Fetal Heart	0.0
Breast ca. MCF-7	0.0	Heart Pool	0.0
Breast ca. MDA-MB-231	0.0	Lymph Node Pool	0.7
Breast ca. BT 549	0.0	Fetal Skeletal Muscle	0.0
Breast ca. T47D	0.0	Skeletal Muscle Pool	0.0
Breast ca. MDA-N	0.0	Spleen Pool	0.0
Breast Pool	2.3	Thymus Pool	0.0
Trachea	0.0	CNS cancer (glio/astro) U87-MG	0.0
Lung	0.0	CNS cancer (glio/astro) U-118-MG	0.0
Fetal Lung	0.0	CNS cancer (neuro;met) SK-N-AS	0.0
Lung ca. NCI-N417	0.0	CNS cancer (astro) SF-539	0.0
Lung ca. LX-1	0.0	CNS cancer (astro) SNB-75	0.0
Lung ca. NCI-H146	0.0	CNS cancer (glio) SNB-19	0.0
Lung ca. SHP-77	0.0	CNS cancer (glio) SF-295	0.0
Lung ca. A549	0.0	Brain (Amygdala) Pool	75.8
Lung ca. NCI-H526	0.0	Brain (cerebellum)	100.0
Lung ca. NCI-H23	0.0	Brain (fetal)	69.3
Lung ca. NCI-H460	0.2	Brain (Hippocampus) Pool	53.2
Lung ca. HOP-62	0.0	Cerebral Cortex Pool	72.2

Lung ca. NCI-H522	0.0	Brain (Substantia nigra) Pool	80.7
Liver	0.0	Brain (Thalamus) Pool	96.6
Fetal Liver	0.0	Brain (whole)	78.5
Liver ca. HepG2	0.0	Spinal Cord Pool	25.0
Kidney Pool	0.0	Adrenal Gland	4.3
Fetal Kidney	0.5	Pituitary gland Pool	0.0
Renal ca. 786-0	0.0	Salivary Gland	0.0
Renal ca. A498	0.0	Thyroid (female)	0.0
Renal ca. ACHN	0.0	Pancreatic ca. CAPAN2	0.0
Renal ca. UO-31	0.0	Pancreas Pool	0.0

CNS_neurodegeneration_v1.0 Summary: Ag5220 This panel confirms the expression of this gene at low levels in the brains of an independent group of individuals: However, no differential expression of this gene was detected between Alzheimer's diseased postmortem brains and those of non-demented controls in this experiment. Please see Panel 1.5 for a discussion of this gene in treatment of central nervous system disorders.

General_screening_panel_v1.5 Summary: Ag5220 Highest expression of this gene is deted in cerebellum (CT=27). High expression of this gene is mainly seen in all the region of central nervous system examined, including amygdala, hippocampus, substantia nigra, thalamus, cerebellum, cerebral cortex, and spinal cord. Therefore, therapeutic modulation of this gene product may be useful in the treatment of central nervous system disorders such as Alzheimer's disease, Parkinson's disease, epilepsy, multiple sclerosis, schizophrenia and depression.

Panel 4.1D Summary: Ag5220 Expression of this gene is low/undetectable (CTs > 35) across all of the samples on this panel.

AI. CG151297-01: CALMODULIN-DEPENDENT PHOSPHODIESTERASE.

Expression of gene CG151297-01 was assessed using the primer-probe set Ag7165, described in Table AIA. Results of the RTQ-PCR runs are shown in Table AIB. Please note that CG151297-01 represents a full-length physical clone.

<u>Table AIA. Probe Name Ag7165</u>

Primers	Sequencs	Length	Start Position	SEQ ID No

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Forward	5'-agaatgtaccgaaaaacattttctct-3'	26	481	387
Probe	TET-5'-ttcctcttatagaggaagcctcaaaag ccg-3'-TAMRA	30	536	388
Reverse	5'-tgcttgccacataggaagaa-3'	20	570	389

Table AIB. Panel 4.1D

Tissue Name	Rel. Ex.(%) Ag7165, Run 307719896	Tissue Name	Rel. Exp.(%) Ag7165, Run 307719896
Secondary Th1 act	0.0	HUVEC IL-1beta	0.0
Secondary Th2 act	0.0	HUVEC IFN gamma	0.0
Secondary Tr1 act	0.0	HUVEC TNF alpha + IFN gamma	0.0
Secondary Th1 rest	0.0	HUVEC TNF alpha + IL4	0.0
Secondary Th2 rest	0.0	HUVEC IL-11	0.0
Secondary Tr1 rest	0.0	Lung Microvascular EC none	0.0
Primary Th1 act	0.0	Lung Microvascular EC TNFalpha + IL-1beta	0.0
Primary Th2 act	0.0	Microvascular Dermal EC none	0.0
Primary Tr1 act	0.0	Microsvasular Dermal EC TNFalpha + IL-1beta	0.0
Primary Th1 rest	0.0	Bronchial epithelium TNFalpha + IL1beta	0.0
Primary Th2 rest	0.0	Small airway epithelium none	0.0
Primary Tr1 rest	0.0	Small airway epithelium TNFalpha + IL-1beta	0.0
CD45RA CD4 lymphocyte act	0.0	Coronery artery SMC rest	0.0
CD45RO CD4 lymphocyte act	0.0	Coronery artery SMC TNFalpha + IL-1beta	0.0
CD8 lymphocyte act	0.0	Astrocytes rest	0.0
Secondary CD8 lymphocyte rest	0.0	Astrocytes TNFalpha + IL-1beta	0.0
Secondary CD8 lymphocyte act	0.0	KU-812 (Basophil) rest	0.0
CD4 lymphocyte none	0.0	KU-812 (Basophil) PMA/ionomycin	0.0
2ry Th1/Th2/Tr1_anti-CD95 CH11	0.0	CCD1106 (Keratinocytes) none	0.0
LAK cells rest	0.0	CCD1106 (Keratinocytes) TNFalpha + IL-1beta	0.0
LAK cells IL-2	0.0	Liver cirrhosis	100.0
LAK cells IL-2+IL-12	0.0	NCI-H292 none	0.0
LAK cells IL-2+IFN gamma	0.0	NCI-H292 IL-4	0.0
LAK cells IL-2+ IL-18	0.0	NCI-H292 IL-9	0.0
LAK cells PMA/ionomycin	0.0	NCI-H292 IL-13	0.0

NK Cells IL-2 rest	0.0	NCI-H292 IFN gamma	0.0
Two Way MLR 3 day	0.0	HPAEC none	0.0
Two Way MLR 5 day	0.0	HPAEC TNF alpha + IL-1 beta	0.0
Two Way MLR 7 day	0.0	Lung fibroblast none	0.0
PBMC rest	0.0	Lung fibroblast TNF alpha + IL-1 beta	0.0
PBMC PWM	0.0	Lung fibroblast IL-4	0.0
PBMC PHA-L	0.0	Lung fibroblast IL-9	0.0
Ramos (B cell) none	0.0	Lung fibroblast IL-13	0.0
Ramos (B cell) ionomycin	0.0	Lung fibroblast IFN gamma	0.0
B lymphocytes PWM	0.0	Dermal fibroblast CCD1070 rest	0.0
B lymphocytes CD40L and IL-4	0.0	Dermal fibroblast CCD1070 TNF alpha	0.0
EOL-1 dbcAMP	0.0	Dermal fibroblast CCD1070 IL-1 beta	0.0
EOL-1 dbcAMP PMA/ionomycin	0.0	Dermal fibroblast IFN gamma	0.0
Dendritic cells none	0.0	Dermal fibroblast IL-4	0.0
Dendritic cells LPS	0.0	Dermal Fibroblasts rest	0.0
Dendritic cells anti-CD40	0.0	Neutrophils TNFa+LPS	0.0
Monocytes rest	0.0	Neutrophils rest	0.0
Monocytes LPS	0.0	Colon	0.0
Macrophages rest	0.0	Lung	0.0
Macrophages LPS	0.0	Thymus	0.0
HUVEC none	0.0	Kidney	0.0
HUVEC starved	0.0		

CNS_neurodegeneration_v1.0 Summary: Ag7165 Expression of this gene is low/undetectable (CTs > 35) across all of the samples on this panel.

Panel 4.1D Summary: Ag7165 Moderate level of expression of this gene is detected mainly in the liver cirrhosis sample (CT=31.5). The presence of this gene in liver cirrhosis (a component of which involves liver inflammation and fibrosis) suggests that antibodies to the protein encoded by this gene could also be used for the diagnosis of liver cirrhosis. Furthermore, therapeutic agents involving this gene may be useful in reducing or inhibiting the inflammation associated with fibrotic and inflammatory diseases.

AJ. CG152256-01: Phosphatidylserine synthase.

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Expression of gene CG152256-01 was assessed using the primer-probe set Ag6718, described in Table AJA. Results of the RTQ-PCR runs are shown in Tables AJB, AJC and AJD.

Table AJA. Probe Name Ag6718

Primers		Length	Start Position	SEQ ID No
Forward	5'-gagcctcgcttccgattat-3'	19	2012	390
Probe	TET-5'-tcccttcccaatattattcatccaga-3'-TAMRA	26	2031	391
Reverse	5'-ctctagcaggtttgcttttgtg-3'	22	2070	392

Table AJB. CNS_neurodegeneration_v1.0

Tissue Name	Rel. Exp.(%) Ag6718, Run 276596848	issue Name	Rel. Exp.(%) Ag6718, Run 276596848
AD 1 Hippo	19.8	Control (Path) 3 Temporal Ctx	2.6
AD 2 Hippo	26.6	Control (Path) 4 Temporal Ctx	15.3
AD 3 Hippo	4.3	AD 1 Occipital Ctx	9.9
AD 4 Hippo	3.7	AD 2 Occipital Ctx (Missing)	0.0
AD 5 Hippo	58.6	AD 3 Occipital Ctx	7.1
AD 6 Hippo	45.4	AD 4 Occipital Ctx	15.9
Control 2 Hippo	28.5	AD 5 Occipital Ctx	26.6
Control 4 Hippo	8.4	AD 6 Occipital Ctx	15.1
Control (Path) 3 Hippo	3.1	Control 1 Occipital Ctx	3.6
AD 1 Temporal Ctx	4.8	Control 2 Occipital Ctx	67.4
AD 2 Temporal Ctx	24.7	Control 3 Occipital Ctx	31.2
AD 3 Temporal Ctx	7.5	Control 4 Occipital Ctx	1.8
AD 4 Temporal Ctx	10.5	Control (Path) 1 Occipital Ctx	100.0
AD 5 Inf Temporal Ctx	62.9	Control (Path) 2 Occipital Ctx	9.5
AD 5 Sup Temporal Ctx	46.3	Control (Path) 3 Occipital Ctx	5.3
AD 6 Inf Temporal Ctx	43.5	Control (Path) 4 Occipital Ctx	10.0
AD 6 Sup Temporal Ctx	43.2	Control 1 Parietal Ctx	3.8
Control 1 Temporal Ctx	4.1	Control 2 Parietal Ctx	27.9
Control 2 Temporal Ctx	59.0	Control 3 Parietal Ctx	15.0
Control 3 Temporal Ctx	17.6	Control (Path) 1 Parietal Ctx	89.5
Control 3 Temporal Ctx	5.0	Control (Path) 2 Parietal Ctx	10.2
Control (Path) 1 Temporal Ctx	57.0	Control (Path) 3 Parietal Ctx	7.0
Control (Path) 2 Temporal Ctx	30.4	Control (Path) 4 Parietal Ctx	27.9

Table AJC. General screening panel v1.6

5

Tissue Name	Rel. Exp.(%) Ag6718, Run 277223813	issue Name	Rel. Exp.(%) Ag6718, Run 277223813
Adipose	2.3	Renal ca. TK-10	34.4
Melanoma* Hs688(A).T	16.4	Bladder	22.2
Melanoma* Hs688(B).T	20.0	Gastric ca. (liver met.) NCI-N87	54.0
Melanoma* M14	30.6	Gastric ca. KATO III	48.3
Melanoma* LOXIMVI	55.1	Colon ca. SW-948	31.0
Melanoma* SK-MEL-5	81.8	Colon ca. SW480	87.1
Squamous cell carcinoma SCC-4	23.5	Colon ca.* (SW480 met) SW620	69.7
Testis Pool	5.2	Colon ca. HT29	0.0
Prostate ca.* (bone met) PC-3	**100.0**	Colon ca. HCT-116	51.4
Prostate Pool	1.8	Colon ca. CaCo-2	15.9
Placenta	2.6	Colon cancer tissue	23.5
Uterus Pool	0.8	Colon ca. SW1116	25.0
Ovarian ca. OVCAR-3	27.4	Colon ca. Colo-205	21.9
Ovarian ca. SK-OV-3	29.9	Colon ca. SW-48	24.1
Ovarian ca. OVCAR-4	33.0	Colon Pool	12.4
Ovarian ca. OVCAR-5	59.9	Small Intestine Pool	4.8
Ovarian ca. IGROV-1	47.6	Stomach Pool	1.8
Ovarian ca. OVCAR-8	32.8	Bone Marrow Pool	0.0
Ovary	11.7	Fetal Heart	14.2
Breast ca. MCF-7	18.9	Heart Pool	11.6
Breast ca. MDA-MB-231	48.0	Lymph Node Pool	3.8
Breast ca. BT 549	31.6	Fetal Skeletal Muscle	3.3
Breast ca. T47D	3.6	Skeletal Muscle Pool	0.0
Breast ca. MDA-N	17.9	Spleen Pool	2.0
Breast Pool	7.0	Thymus Pool	11.7
Trachea	9.2	CNS cancer (glio/astro) U87-MG	32.3
Lung	2.4	CNS cancer (glio/astro) U-118-MG	43.2
Fetal Lung	4.9	CNS cancer (neuro;met) SK-N-AS	25.9
Lung ca. NCI-N417	15.0	CNS cancer (astro) SF-539	29.5
Lung ca. LX-1	17.6	CNS cancer (astro) SNB-75	59.0
Lung ca. NCI-H146	23.7	CNS cancer (glio) SNB-19	29.7
Lung ca. SHP-77	53.2	CNS cancer (glio) SF-295	59.5
Lung ca. A549	28.3	Brain (Amygdala) Pool	10.4
Lung ca. NCI-H526	24.3	Brain (cerebellum)	34.4
Lung ca. NCI-H23	71.7	Brain (fetal)	17.3

Lung ca. NCI-H460	14.2	Brain (Hippocampus) Pool	9.4
Lung ca. HOP-62	32.3	Cerebral Cortex Pool	7.4
Lung ca. NCI-H522	16.4	Brain (Substantia nigra) Pool	3.9
Liver	1.0	Brain (Thalamus) Pool	6.9
Fetal Liver	2.3	Brain (whole)	6.5
Liver ca. HepG2	19.2	Spinal Cord Pool	5.6
Kidney Pool	15.2	Adrenal Gland	10.3
Fetal Kidney	4.1	Pituitary gland Pool	1.1
Renal ca. 786-0	61.6	Salivary Gland	3.2
Renal ca. A498	5.6	Thyroid (female)	11.5
Renal ca. ACHN	24.7	Pancreatic ca. CAPAN2	28.1
Renal ca. UO-31	33.9	Pancreas Pool	8.3

Table AJD. Panel 4.1D

5

Tissue Name	Rel. Ex.(%) Ag6718, Run 276596888	Tissue Name	Rel. Exp.(%) Ag6718, Run 276596888
Secondary Th1 act	51.4	HUVEC IL-1beta	18.0
Secondary Th2 act	39.5	HUVEC IFN gamma	16.5
Secondary Tr1 act	19.3	HUVEC TNF alpha + IFN gamma	4.5
Secondary Th1 rest	5.3	HUVEC TNF alpha + IL4	3.1
Secondary Th2 rest	4.5	HUVEC IL-11	0.0
Secondary Tr1 rest	5.9	Lung Microvascular EC none	13.9
Primary Th1 act	3.5	Lung Microvascular EC TNFalpha + IL-1beta	0.7
Primary Th2 act	20.7	Microvascular Dermal EC none	3.0
Primary Tr1 act	12.8	Microsvasular Dermal EC TNFalpha + IL-1beta	1.2
Primary Th1 rest	1.6	Bronchial epithelium TNFalpha + IL1beta	5.8
Primary Th2 rest	5.8	Small airway epithelium none	6.3
Primary Tr1 rest	0.7	Small airway epithelium TNFalpha + IL-1beta	9.7
CD45RA CD4 lymphocyte act	26.4	Coronery artery SMC rest	7.1
CD45RO CD4 lymphocyte act	30.8	Coronery artery SMC TNFalpha + IL-1beta	8.4
CD8 lymphocyte act	7.6	Astrocytes rest	3.3
Secondary CD8 lymphocyte rest	6.3	Astrocytes TNFalpha + IL-1beta	2.9
Secondary CD8 lymphocyte act	1.5	KU-812 (Basophil) rest	44.8
CD4 lymphocyte none	3.6	KU-812 (Basophil) PMA/ionomycin	28.1

2ry Th1/Th2/Tr1_anti-CD95 CH11	2.9	CCD1106 (Keratinocytes) none	27.5
LAK cells rest	4.5	CCD1106 (Keratinocytes) TNFalpha + IL-1beta	5.1
LAK cells IL-2	9.9	Liver cirrhosis	0.8
LAK cells IL-2+IL-12	0.7	NCI-H292 none	8.0
LAK cells IL-2+IFN gamma	4.2	NCI-H292 IL-4	10.2
LAK cells IL-2+ IL-18	1.4	NCI-H292 IL-9	19.2
LAK cells PMA/ionomycin	18.7	NCI-H292 IL-13	14.8
NK Cells IL-2 rest	21.0	NCI-H292 IFN gamma	6.8
Two Way MLR 3 day	7.6	HPAEC none	3.7
Two Way MLR 5 day	5.2	HPAEC TNF alpha + IL-1 beta	8.5
Two Way MLR 7 day	4.3	Lung fibroblast none	6.8
PBMC rest	1.4	Lung fibroblast TNF alpha + IL-1 beta	1.9
PBMC PWM	3.0	Lung fibroblast IL-4	6.1
PBMC PHA-L	4.1	Lung fibroblast IL-9	10.0
Ramos (B cell) none	42.9	Lung fibroblast IL-13	7.7
Ramos (B cell) ionomycin	22.1	Lung fibroblast IFN gamma	16.4
B lymphocytes PWM	10.8	Dermal fibroblast CCD1070 rest	33.9
B lymphocytes CD40L and IL-4	12.2	Dermal fibroblast CCD1070 TNF alpha	100.0
EOL-1 dbcAMP	39.0	Dermal fibroblast CCD1070 IL-1 beta	17.4
EOL-1 dbcAMP PMA/ionomycin	14.1	Dermal fibroblast IFN gamma	6.7
Dendritic cells none	13.5	Dermal fibroblast IL-4	10.4
Dendritic cells LPS	2.5	Dermal Fibroblasts rest	6.9
Dendritic cells anti-CD40	4.5	Neutrophils TNFa+LPS	0.4
Monocytes rest	0.6	Neutrophils rest	0.7
Monocytes LPS	3.9	Colon	0.8
Macrophages rest	1.4	Lung	0.6
Macrophages LPS	3.8	Thymus	2.9
HUVEC none	11.1	Kidney	8.1
HUVEC starved	6.4		

CNS_neurodegeneration_v1.0 Summary: Ag6718 This panel confirms the expression of this gene at low levels in the brains of an independent group of individuals. However, no differential expression of this gene was detected between Alzheimer's diseased postmortem brains and those of non-demented controls in this experiment. Please see Panel 1.6 for a discussion of this gene in treatment of central nervous system disorders.

General_screening_panel_v1.6 Summary: Ag6718 Highest expression of this gene is detected in prostate cancer PC3 cell line (CT=31.9). Moderate levels of expression of this gene is also seen in cluster of cancer cell lines derived from pancreatic, gastric, colon, lung, liver, renal, breast, ovarian, prostate, squamous cell carcinoma, melanoma and brain cancers. Thus, expression of this gene could be used as a marker to detect the presence of these cancers. Furthermore, therapeutic modulation of the expression or function of this gene may be effective in the treatment of pancreatic, gastric, colon, lung, liver, renal, breast, ovarian, prostate, squamous cell carcinoma, melanoma and brain cancers.

In addition, this gene is expressed at low levels in cerebellum and fetal brain. Therefore, therapeutic modulation of this gene product may be useful in the treatment of central nervous system disorders such as ataxia and autism.

Panel 4.1D Summary: Ag6718 Highest expression of this gene is detected in TNF alpha treated dermal fibroblasts (CT=32). Moderate to low levels of expression of this gene is detected in activated polarized, naive and memory T cells, PMA/ionomycin treated LAK cells, resting IL-2 treated NK cells, Ramos B cells, eosinophils, activated HUVEC cells, lung microvascular endothelial cells, basophils and activated mucoepidermoid NCI-H292 cells. Therefore, therapeutic modulation of this gene or its protein product may lead to the alteration of functions associated with these cell types and lead to improvement of the symptoms of patients suffering from autoimmune and inflammatory diseases such as asthma, allergies, inflammatory bowel disease, lupus erythematosus, psoriasis, rheumatoid arthritis, and osteoarthritis.

AK. CG173017-01: RETINOIC ACID RECEPTOR RXR-BETA.

Expression of gene CG173017-01 was assessed using the primer-probe set Ag7565, described in Table AKA.

Table AKA. Probe Name Ag7565

Primers		Length	Start Position	SEQ ID No
Forward	5'-ctggacgggacgggat-3'	16	222	393
Probe	TET-5'-acatagccgtttgccagccccag-3'-TAMRA	23	261	394

Reverse	5'-cttctgtcccgcagatt-3'	18	286	395

CNS_neurodegeneration_v1.0 Summary: Ag7565 Expression of this gene is low/undetectable in all samples on this panel (CTs>35).

Panel 4.1D Summary: Ag7565 Expression of this gene is low/undetectable in all samples on this panel (CTs>35).

AL. CG173347-01: Novel Serum paraoxonase/arylesterase 3.

Expression of gene CG173347-01 was assessed using the primer-probe set Ag7564, described in Table ALA.

Table ALA. Probe Name Ag7564

Primers	Sequence	Length	Start Position	SEQ ID No
Forward	5'-gaaagtggctctgaagatattgatatact-3'	29	153	396
Probe	TET-5'-tcctagtgggctggcttttatctcc-3'-TAMRA	25	182	397
Reverse	5'-actccaacagacctgcagact-3'	21	207	398

CNS_neurodegeneration_v1.0 Summary: Ag7564 Expression of this gene is low/undetectable in all samples on this panel (CTs>35).

Panel 4.1D Summary: Ag7564 Expression of this gene is low/undetectable in all samples on this panel (CTs>35).

AM. CG56234-02: Splice variant of PCK2.

Expression of gene CG56234-02 was assessed using the primer-probe set Ag5111, described in Table AMA. Results of the RTQ-PCR runs are shown in Tables AMB, AMC, AMD and AME.

Table AMA. Probe Name Ag5111

Primers		Length	Start Position	SEQ ID No
Forward	5'-ctgggaggccccaga-3'	15	1377	399

| Probe | TET-5'-tgtccccattgacgccatcatc-3'-TAMRA | 22 | 1395 | 400 |
| Reverse | 5'-gatgatcttccctttgggtct-3' | 21 | 1429 | 401 |

Table AMB. General screening panel v1.5

Tissue Name	Rel. Exp.(%) Ag5111, Run 228980587	issue Name	Rel. Exp.(%) Ag5111, Run 228980587
Adipose	2.0	Renal ca. TK-10	29.1
Melanoma* Hs688(A).T	31.9	Bladder	12.1
Melanoma* Hs688(B).T	28.3	Gastric ca. (liver met.) NCI-N87	31.4
Melanoma* M14	9.9	Gastric ca. KATO III	28.1
Melanoma* LOXIMVI	4.5	Colon ca. SW-948	17.9
Melanoma* SK-MEL-5	39.8	Colon ca. SW480	14.9
Squamous cell carcinoma SCC-4	4.7	Colon ca.* (SW480 met) SW620	29.5
Testis Pool	1.6	Colon ca. HT29	8.6
Prostate ca.* (bone met) PC-3	55.1	Colon ca. HCT-116	11.0
Prostate Pool	0.5	Colon ca. CaCo-2	44.4
Placenta	0.3	Colon cancer tissue	9.7
Uterus Pool	0.6	Colon ca. SW1116	1.4
Ovarian ca. OVCAR-3	13.6	Colon ca. Colo-205	6.6
Ovarian ca. SK-OV-3	5.3	Colon ca. SW-48	14.4
Ovarian ca. OVCAR-4	7.1	Colon Pool	0.1
Ovarian ca. OVCAR-5	34.6	Small Intestine Pool	0.6
Ovarian ca. IGROV-1	22.5	Stomach Pool	1.1
Ovarian ca. OVCAR-8	100.0	Bone Marrow Pool	0.5
Ovary	0.0	Fetal Heart	0.0
Breast ca. MCF-7	87.7	Heart Pool	0.0
Breast ca. MDA-MB-231	12.6	Lymph Node Pool	0.8
Breast ca. BT 549	75.8	Fetal Skeletal Muscle	0.6
Breast ca. T47D	10.1	Skeletal Muscle Pool	0.4
Breast ca. MDA-N	16.4	Spleen Pool	1.7
Breast Pool	0.5	Thymus Pool	0.4
Trachea	4.3	CNS cancer (glio/astro) U87-MG	18.8
Lung	0.0	CNS cancer (glio/astro) U-118-MG	9.3
Fetal Lung	2.0	CNS cancer (neuro;met) SK-N-AS	7.5
Lung ca. NCI-N417	1.8	CNS cancer (astro) SF-539	11.3
Lung ca. LX-1	8.2	CNS cancer (astro) SNB-75	48.6
Lung ca. NCI-H146	11.1	CNS cancer (glio) SNB-19	31.0
Lung ca. SHP-77	11.3	CNS cancer (glio) SF-295	32.5

Lung ca. A549	11.4	Brain (Amygdala) Pool	0.4
Lung ca. NCI-H526	1.8	Brain (cerebellum)	0.3
Lung ca. NCI-H23	83.5	Brain (fetal)	0.3
Lung ca. NCI-H460	27.0	Brain (Hippocampus) Pool	2.5
Lung ca. HOP-62	1.0	Cerebral Cortex Pool	0.4
Lung ca. NCI-H522	67.4	Brain (Substantia nigra) Pool	0.0
Liver	6.3	Brain (Thalamus) Pool	1.0
Fetal Liver	6.7	Brain (whole)	0.7
Liver ca. HepG2	24.7	Spinal Cord Pool	1.1
Kidney Pool	0.8	Adrenal Gland	1.6
Fetal Kidney	1.0	Pituitary gland Pool	0.4
Renal ca. 786-0	8.7	Salivary Gland	0.9
Renal ca. A498	1.5	Thyroid (female)	0.7
Renal ca. ACHN	9.3	Pancreatic ca. CAPAN2	12.8
Renal ca. UO-31	1.9	Pancreas Pool	0.8

Table AMC. General_screening_panel_v1.6

5

Tissue Name	Rel. Exp.(%) Ag5111, Run 277218717	Rel. Exp.(%) Ag5111, Run 277731246	Rel. Exp.() Ag5111, Run 278368614	Tissue Name	Rel. Exp.(%) Ag5111, Run 277218717	Rel. Exp.(%) Ag5111, Run 277731246	Rel. Exp.(%) Ag5111, Run 278368614
Adipose	0.5	0.0	1.5	Renal ca. TK-10	24.7	20.2	33.0
Melanoma* Hs688(A).T	26.1	29.5	31.6	Bladder	6.7	6.1	11.6
Melanoma* Hs688(B).T	25.2	32.1	31.9	Gastric ca. (liver met.) NCI-N87	21.3	22.5	36.1
Melanoma* M14	5.6	9.7	7.5	Gastric ca. KATO III	14.6	12.2	19.2
Melanoma* LOXIMVI	3.0	0.0	4.2	Colon ca. SW-948	18.8	16.5	23.5
Melanoma* SK-MEL-5	28.7	57.0	39.8	Colon ca. SW480	11.8	7.3	19.5
Squamous cell carcinoma SCC-4	4.8	4.2	5.1	Colon ca.* (SW480 met) SW620	23.0	19.9	35.6
Testis Pool	2.0	0.0	1.4	Colon ca. HT29	10.2	4.2	8.2
Prostate ca.* (bone met) PC-3	33.2	44.4	57.8	Colon ca. HCT-116	9.6	7.6	19.9

Prostate Pool	0.3	0.0	0.6	Colon ca. CaCo-2	9.4	25.0	36.9
Placenta	0.3	0.0	1.1	Colon cancer tissue	6.0	0.0	6.6
Uterus Pool	0.0	0.0	0.6	Colon ca. SW1116	2.3	0.0	1.7
Ovarian ca. OVCAR-3	12.7	8.2	18.2	Colon ca. Colo-205	5.1	4.7	5.9
Ovarian ca. SK-OV-3	5.3	6.5	12.2	Colon ca. SW-48	9.0	0.0	11.6
Ovarian ca. OVCAR-4	4.0	5.2	5.8	Colon Pool	0.7	0.0	0.7
Ovarian ca. OVCAR-5	31.6	24.8	34.2	Small Intestine Pool	0.3	0.0	0.8
Ovarian ca. IGROV-1	19.2	12.8	27.2	Stomach Pool	1.2	0.0	2.3
Ovarian ca. OVCAR-8	**100.0**	**100.0**	**100.0**	Bone Marrow Pool	0.0	0.0	0.0
Ovary	0.0	0.0	0.2	Fetal Heart	0.0	0.0	0.3
Breast ca. MCF-7	54.0	51.4	77.9	Heart Pool	0.4	0.0	0.0
Breast ca. MDA-MB-231	8.5	7.6	7.7	Lymph Node Pool	1.2	0.0	0.0
Breast ca. BT 549	47.0	30.4	49.0	Fetal Skeletal Muscle	0.0	0.0	0.0
Breast ca. T47D	5.1	6.5	7.1	Skeletal Muscle Pool	0.0	0.0	0.0
Breast ca. MDA-N	6.1	6.0	24.5	Spleen Pool	0.7	0.0	2.5
Breast Pool	0.3	0.0	0.3	Thymus Pool	0.5	0.0	1.8
Trachea	3.3	0.0	8.3	CNS cancer (glio/astro) U87-MG	12.9	7.9	13.8
Lung	0.0	0.0	0.0	CNS cancer (glio/astro) U-118-MG	5.9	4.4	8.1
Fetal Lung	0.9	0.0	2.1	CNS cancer (neuro;met) SK-N-AS	6.4	4.9	6.7
Lung ca. NCI-N417	1.3	0.0	3.8	CNS cancer (astro) SF-539	5.8	6.4	8.5
Lung ca. LX-1	5.5	7.8	9.5	CNS cancer (astro) SNB-75	25.0	29.9	26.8
Lung ca. NCI-H146	8.0	8.5	11.5	CNS cancer (glio) SNB-19	23.8	20.7	29.5
Lung ca. SHP-77	12.2	14.3	21.3	CNS cancer (glio) SF-295	38.2	28.7	46.7

Lung ca. A549	11.5	11.7	15.9	Brain (Amygdala) Pool	0.8	0.0	1.1
Lung ca. NCI-H526	1.8	0.0	1.7	Brain (cerebellum)	1.0	0.0	1.1
Lung ca. NCI-H23	42.6	68.8	55.1	Brain (fetal)	0.0	0.0	0.4
Lung ca. NCI-H460	16.7	23.5	38.4	Brain (Hippocampus) Pool	0.4	0.0	1.2
Lung ca. HOP-62	2.0	0.0	3.0	Cerebral Cortex Pool	0.0	0.0	0.6
Lung ca. NCI-H522	41.5	64.2	87.1	Brain (Substantia nigra) Pool	0.0	0.0	0.4
Liver	4.4	4.6	7.1	Brain (Thalamus) Pool	0.0	0.0	0.0
Fetal Liver	5.8	3.3	8.7	Brain (whole)	6.7	0.0	2.8
Liver ca. HepG2	15.7	16.3	18.8	Spinal Cord Pool	0.6	0.0	0.5
Kidney Pool	0.7	0.0	0.3	Adrenal Gland	1.4	0.0	1.4
Fetal Kidney	0.9	0.0	1.0	Pituitary gland Pool	0.0	0.0	0.7
Renal ca. 786-0	9.3	8.1	13.8	Salivary Gland	0.8	0.0	1.8
Renal ca. A498	1.1	0.0	2.0	Thyroid (female)	1.0	0.0	2.1
Renal ca. ACHN	5.8	6.0	10.8	Pancreatic ca. CAPAN2	13.1	9.6	19.9
Renal ca. UO-31	2.4	0.0	3.3	Pancreas Pool	4.8	0.0	7.3

Table AMD. Panel 4.1D

5

Tissue Name	Rel. Exp.(%) g5111, Run 226444761	Rel. Exp.(%) Ag5111, Run 276596864	Tissue Name	Rel. Exp.(%) Ag5111, Run 226444761	Rel. Exp.(%) Ag5111, Run 276596864
Secondary Th1 act	90.8	58.6	HUVEC IL-1beta	18.7	10.7
Secondary Th2 act	40.9	57.8	HUVEC IFN gamma	2.8	6.2
Secondary Tr1 act	57.4	16.5	HUVEC TNF alpha + IFN gamma	5.0	6.2
Secondary Th1 rest	27.2	8.4	HUVEC TNF alpha + IL4	23.2	8.8
Secondary Th2 rest	6.0	0.0	HUVEC IL-11	2.3	0.0

Secondary Tr1 rest	7.2	4.0	Lung Microvascular EC none	3.2	15.4
Primary Th1 act	32.8	5.0	Lung Microvascular EC TNFalpha + IL-1beta	6.4	0.0
Primary Th2 act	49.0	19.9	Microvascular Dermal EC none	6.6	0.0
Primary Tr1 act	50.0	38.4	Microsvasular Dermal EC TNFalpha + IL-1beta	0.0	0.0
Primary Th1 rest	6.0	8.5	Bronchial epithelium TNFalpha + IL1beta	8.7	6.9
Primary Th2 rest	6.4	6.3	Small airway epithelium none	2.2	0.0
Primary Tr1 rest	18.0	0.0	Small airway epithelium TNFalpha + IL-1beta	11.8	0.0
CD45RA CD4 lymphocyte act	95.9	76.8	Coronery artery SMC rest	18.3	10.2
CD45RO CD4 lymphocyte act	95.3	**100.0**	Coronery artery SMC TNFalpha + IL-1beta	9.4	8.8
CD8 lymphocyte act	77.4	4.5	Astrocytes rest	2.1	0.0
Secondary CD8 lymphocyte rest	90.1	17.3	Astrocytes TNFalpha + IL-1beta	0.0	0.0
Secondary CD8 lymphocyte act	21.0	7.7	KU-812 (Basophil) rest	25.9	10.2
CD4 lymphocyte none	0.0	0.0	KU-812 (Basophil) PMA/ionomycin	26.8	21.2
2ry Th1/Th2/Tr1_anti-CD95 CH11	5.4	0.0	CCD1106 (Keratinocytes) none	15.2	4.9
LAK cells rest	43.5	19.9	CCD1106 (Keratinocytes) TNFalpha + IL-1beta	9.0	12.3
LAK cells IL-2	52.1	18.4	Liver cirrhosis	8.3	0.0
LAK cells IL-2+IL-12	33.7	0.0	NCI-H292 none	15.3	3.4
LAK cells IL-2+IFN gamma	57.0	6.6	NCI-H292 IL-4	13.5	17.2
LAK cells IL-2+ IL-18	46.0	9.5	NCI-H292 IL-9	14.2	14.1
LAK cells PMA/ionomycin	43.5	24.5	NCI-H292 IL-13	29.1	11.3
NK Cells IL-2 rest	60.7	37.4	NCI-H292 IFN gamma	44.8	7.2
Two Way MLR 3 day	32.1	10.3	HPAEC none	2.0	0.0
Two Way MLR 5 day	53.2	3.6	HPAEC TNF alpha + IL-1 beta	7.2	7.0
Two Way MLR 7 day	23.5	9.6	Lung fibroblast none	21.2	15.9

PBMC rest	6.1	0.0	Lung fibroblast TNF alpha + IL-1 beta	11.5	0.0
PBMC PWM	23.5	9.1	Lung fibroblast IL-4	2.4	0.0
PBMC PHA-L	35.8	12.2	Lung fibroblast IL-9	17.6	5.4
Ramos (B cell) none	58.6	16.7	Lung fibroblast IL-13	13.4	0.0
Ramos (B cell) ionomycin	71.7	92.7	Lung fibroblast IFN gamma	11.6	3.1
B lymphocytes PWM	21.6	14.8	Dermal fibroblast CCD1070 rest	99.3	64.6
B lymphocytes CD40L and IL-4	29.7	23.2	Dermal fibroblast CCD1070 TNF alpha	74.7	88.9
EOL-1 dbcAMP	32.3	32.8	Dermal fibroblast CCD1070 IL-1 beta	29.9	50.0
EOL-1 dbcAMP PMA/ionomycin	10.6	3.2	Dermal fibroblast IFN gamma	13.3	0.0
Dendritic cells none	66.0	24.5	Dermal fibroblast IL-4	12.2	0.0
Dendritic cells LPS	31.4	0.0	Dermal Fibroblasts rest	0.0	0.0
Dendritic cells anti-CD40	48.3	28.1	Neutrophils TNFa+LPS	0.0	0.0
Monocytes rest	29.1	0.0	Neutrophils rest	0.0	0.0
Monocytes LPS	37.6	18.0	Colon	32.3	8.2
Macrophages rest	**100.0**	12.9	Lung	3.5	0.0
Macrophages LPS	28.1	16.2	Thymus	12.1	0.0
HUVEC none	7.9	5.7	Kidney	83.5	31.9
HUVEC starved	17.4	8.4			

Table AME. general oncology screening panel_v_2.4

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Tissue Name	Rel. Exp.(%) Ag5111, Run 260280403	Tissue Nme	Rel. Exp.(%) Ag5111, Run 260280403
Colon cancer 1	49.0	Bladder cancer NAT 2	0.0
Colon cancer NAT 1	2.5	Bladder cancer NAT 3	0.0
Colon cancer 2	11.7	Bladder cancer NAT 4	0.0
Colon cancer NAT 2	28.5	Prostate adenocarcinoma 1	5.0
Colon cancer 3	43.5	Prostate adenocarcinoma 2	0.0
Colon cancer NAT 3	53.2	Prostate adenocarcinoma 3	0.0
Colon malignant cancer 4	**100.0**	Prostate adenocarcinoma 4	0.0
Colon normal adjacent tissue 4	8.4	Prostate cancer NAT 5	0.0
Lung cancer 1	12.2	Prostate adenocarcinoma 6	0.0
Lung NAT 1	0.0	Prostate adenocarcinoma 7	0.0

Lung cancer 2	72.2	Prostate adenocarcinoma 8	0.0
Lung NAT 2	0.0	Prostate adenocarcinoma 9	4.0
Squamous cell carcinoma 3	18.8	Prostate cancer NAT 10	0.0
Lung NAT 3	0.0	Kidney cancer 1	7.5
metastatic melanoma 1	0.0	KidneyNAT 1	0.0
Melanoma 2	6.3	Kidney cancer 2	73.2
Melanoma 3	0.0	Kidney NAT 2	9.2
metastatic melanoma 4	0.0	Kidney cancer 3	6.3
metastatic melanoma 5	2.0	Kidney NAT 3	0.0
Bladder cancer 1	0.0	Kidney cancer 4	7.6
Bladder cancer NAT 1	0.0	Kidney NAT 4	84.1
Bladder cancer 2	0.0		

CNS_neurodegeneration_v1.0 Summary: Ag5111 Expression of the CG56234-02 gene is low/undetectable in all samples on this panel (CTs>35).

General_screening_panel_v1.5 Summary: Ag5111 Highest expression of the CG56234-02 gene is seen in an ovarian cancer cell line (CT=30). This gene encodes a splice variant of PEPCK2, the rate-limiting enzyme for gluconeogenesis that has been shown to be regulated in response to hormones and environmental stress. In addition, to the ovarian cancer cell line, this gene is expressed at a moderate level in most of the cancer cell lines used in this panel. Therefore, modulation of the gene product using small molecule drugs may affect the growth and survival of cancer cells. Expression of this gene could potentially be used as a diagnostic marker of the metabolic status of cells and inhibition of activity of this gene prodcut might be used for therapeutic treatment of cancers.

This gene is also moderately expressed (CT values = 34) in adult and fetal liver. Inhibition of this enzyme could potentially decrease hepatic glucose production and thus serve as an effective treatment for Type 2 diabetes, which is characterized by excess hepatic glucose production.

General_screening_panel_v1.6 Summary: Ag5111 Three experiments with the same probe and primer produce results that are in excellent agreement. Highest expression is seen in an ovarian cancer cell line (CTs=31-34) and overall, expression of this gene appears to be more highly associated with cancer cell line samples than with normal tissue samples. These results are also in agreement with results in Panel 1.5. Please see that panel for discussion of this gene.

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Panel 4.1D Summary: Ag 5111 This gene is expressed at low levels in a wide range of cell across this panel (CTs=33.3-34.4), including CD4 T cells (naive and memory T cells), CD8 T cells, B cells and macrophages. Expression of this transcript is also found in dermal fibroblasts and kidney. This transcript encodes a homolog of a key enzyme in glucogenesis and therefore may be important for the metabolic status of all these cell types which contribute to the inflammatory response. Therefore, modulation of the activity or expression of this putative protein by small molecules could affect the activity of these cells and be useful for the treatment of autoimmune diseases such as inflammatory bowel diseases, rheumatoid arthritis, asthma, COPD, psoriasis and lupus.

general oncology screening panel_v_2.4 Summary: Ag5111 Low but significant expression is seen in a colon cancer, a kidney cancer, and a lung cancer (CTs=34-35). This is in agreement with the preferential expression in cancer cell lines seen in Panels 1.5 and 1.6. Please see Panel 1.5 for discussion of this gene in oncology.

AN. CG56836-03: Cathepsin B.

Expression of gene CG56836-03 was assessed using the primer-probe sets Ag2052 and Ag5278, described in Tables ANA, ANB and ANC. Results of the RTQ-PCR runs are shown in Tables AND, ANE, ANF, ANG, ANH, ANI, ANJ and ANK.

Table ANA. Probe Name Ag2052

Primers		Length	Start Position	SEQ ID No
Forward	5'-gtcccaccatcaaagagatca-3'	21	414	402
Probe	TET-5'-agaccagggctcctgtggctcct-3'-TAMRA	23	436	403
Reverse	5'-atgcagatccggtcagagat-3'	20	485	404

Table ANB. Probe Name Ag5277

Primers		Length	Start Position	SEQ ID No
Forward	5'-gatctgcatccacaccaat-3'	19	390	405
Probe	TET-5'-cctgctcacctgcctgctctacaagt-3'-TAMRA	26	441	406
Reverse	5'-cagtcagtgttccaggagtt-3'	20	568	407

Table ANC. Probe Name Ag5278

Primers		Length	Start Position	SEQ ID No
Forward	5'-tatgaatccaatagcgaga-3'	19	653	408
Probe	TET-5'-agctttctctgtgtattcggacttcc-3'-TAMRA	26	715	409
Reverse	5'-tgttggtacactcctgactt-3'	20	749	410

Table AND. AI_comprehensive panel_v1.0

Tissue Name	Rel. Exp.(%) Ag2052, Run 275804031	issue Name	Rel. Exp.(%) Ag2052, Run 275804031
110967 COPD-F	10.2	112427 Match Control Psoriasis-F	15.4
110980 COPD-F	6.4	112418 Psoriasis-M	10.4
110968 COPD-M	12.0	112723 Match Control Psoriasis-M	5.9
110977 COPD-M	14.0	112419 Psoriasis-M	12.9
110989 Emphysema-F	15.6	112424 Match Control Psoriasis-M	4.3
110992 Emphysema-F	20.0	112420 Psoriasis-M	29.7
110993 Emphysema-F	13.8	112425 Match Control Psoriasis-M	14.8
110994 Emphysema-F	6.0	104689 (MF) OA Bone-Backus	29.9
110995 Emphysema-F	33.2	104690 (MF) Adj "Normal" Bone-Backus	15.4
110996 Emphysema-F	8.5	104691 (MF) OA Synovium-Backus	55.9
110997 Asthma-M	6.1	104692 (BA) OA Cartilage-Backus	27.9
111001 Asthma-F	6.7	104694 (BA) OA Bone-Backus	39.5
111002 Asthma-F	11.2	104695 (BA) Adj "Normal" Bone-Backus	23.0
111003 Atopic Asthma-F	9.7	104696 (BA) OA Synovium-Backus	100.0
111004 Atopic Asthma-F	12.2	104700 (SS) OA Bone-Backus	12.2
111005 Atopic Asthma-F	7.4	104701 (SS) Adj "Normal" Bone-Backus	24.3
111006 Atopic Asthma-F	1.7	104702 (SS) OA Synovium-Backus	43.8
111417 Allergy-M	9.0	117093 OA Cartilage Rep7	18.4
112347 Allergy-M	0.0	112672 OA Bone5	17.3
112349 Normal Lung-F	0.0	112673 OA Synovium5	6.6
112357 Normal Lung-F	10.7	112674 OA Synovial Fluid cells5	8.4
112354 Normal Lung-M	3.6	117100 OA Cartilage Rep14	8.4
112374 Crohns-F	10.6	112756 OA Bone9	13.4

112389 Match Control Crohns-F	14.1	112757 OA Synovium9	4.0
112375 Crohns-F	9.9	112758 OA Synovial Fluid Cells9	5.0
112732 Match Control Crohns-F	6.6	117125 RA Cartilage Rep2	19.5
112725 Crohns-M	1.3	113492 Bone2 RA	11.7
112387 Match Control Crohns-M	11.7	113493 Synovium2 RA	3.6
112378 Crohns-M	0.0	113494 Syn Fluid Cells RA	6.7
112390 Match Control Crohns-M	14.5	113499 Cartilage4 RA	6.7
112726 Crohns-M	11.5	113500 Bone4 RA	6.3
112731 Match Control Crohns-M	7.5	113501 Synovium4 RA	5.1
112380 Ulcer Col-F	8.7	113502 Syn Fluid Cells4 RA	3.4
112734 Match Control Ulcer Col-F	15.4	113495 Cartilage3 RA	7.2
112384 Ulcer Col-F	25.7	113496 Bone3 RA	7.0
112737 Match Control Ulcer Col-F	4.1	113497 Synovium3 RA	4.4
112386 Ulcer Col-F	7.1	113498 Syn Fluid Cells3 RA	9.7
112738 Match Control Ulcer Col-F	13.1	117106 Normal Cartilage Rep20	8.1
112381 Ulcer Col-M	0.1	113663 Bone3 Normal	0.0
112735 Match Control Ulcer Col-M	0.4	113664 Synovium3 Normal	0.0
112382 Ulcer Col-M	12.9	113665 Syn Fluid Cells3 Normal	0.0
112394 Match Control Ulcer Col-M	3.3	117107 Normal Cartilage Rep22	3.2
112383 Ulcer Col-M	30.4	113667 Bone4 Normal	6.3
112736 Match Control Ulcer Col-M	11.0	113668 Synovium4 Normal	8.1
112423 Psoriasis-F	5.5	113669 Syn Fluid Cells4 Normal	12.9

Table ANE. General screening panel v1.5

Tissue Name	Rel. Exp.(%) Ag5278, Run 230509757	issue Name	Rel. Exp.(%) Ag5278, Run 230509757
Adipose	0.2	Renal ca. TK-10	6.2
Melanoma* Hs688(A).T	24.0	Bladder	5.1
Melanoma* Hs688(B).T	12.9	Gastric ca. (liver met.) NCI-N87	9.7
Melanoma* M14	51.8	Gastric ca. KATO III	5.7
Melanoma* LOXIMVI	26.6	Colon ca. SW-948	2.1
Melanoma* SK-MEL-5	17.0	Colon ca. SW480	7.0

Squamous cell carcinoma SCC-4	3.2	Colon ca.* (SW480 met) SW620	3.2
Testis Pool	0.5	Colon ca. HT29	0.7
Prostate ca.* (bone met) PC-3	0.6	Colon ca. HCT-116	2.6
Prostate Pool	0.2	Colon ca. CaCo-2	5.3
Placenta	5.4	Colon cancer tissue	14.5
Uterus Pool	0.0	Colon ca. SW1116	2.3
Ovarian ca. OVCAR-3	16.3	Colon ca. Colo-205	7.9
Ovarian ca. SK-OV-3	18.7	Colon ca. SW-48	2.7
Ovarian ca. OVCAR-4	3.9	Colon Pool	1.8
Ovarian ca. OVCAR-5	5.7	Small Intestine Pool	0.7
Ovarian ca. IGROV-1	0.3	Stomach Pool	1.2
Ovarian ca. OVCAR-8	1.3	Bone Marrow Pool	0.3
Ovary	3.2	Fetal Heart	0.5
Breast ca. MCF-7	3.0	Heart Pool	1.2
Breast ca. MDA-MB-231	4.1	Lymph Node Pool	2.9
Breast ca. BT 549	100.0	Fetal Skeletal Muscle	0.3
Breast ca. T47D	2.0	Skeletal Muscle Pool	1.0
Breast ca. MDA-N	1.6	Spleen Pool	2.1
Breast Pool	2.0	Thymus Pool	1.4
Trachea	2.3	CNS cancer (glio/astro) U87-MG	8.1
Lung	0.5	CNS cancer (glio/astro) U-118-MG	12.3
Fetal Lung	2.2	CNS cancer (neuro;met) SK-N-AS	2.0
Lung ca. NCI-N417	0.1	CNS cancer (astro) SF-539	3.4
Lung ca. LX-1	6.1	CNS cancer (astro) SNB-75	27.4
Lung ca. NCI-H146	0.4	CNS cancer (glio) SNB-19	2.4
Lung ca. SHP-77	1.8	CNS cancer (glio) SF-295	26.8
Lung ca. A549	4.1	Brain (Amygdala) Pool	2.1
Lung ca. NCI-H526	0.1	Brain (cerebellum)	6.9
Lung ca. NCI-H23	3.0	Brain (fetal)	1.2
Lung ca. NCI-H460	2.6	Brain (Hippocampus) Pool	1.9
Lung ca. HOP-62	4.0	Cerebral Cortex Pool	3.8
Lung ca. NCI-H522	1.0	Brain (Substantia nigra) Pool	2.6
Liver	1.4	Brain (Thalamus) Pool	2.8
Fetal Liver	10.4	Brain (whole)	5.3
Liver ca. HepG2	8.3	Spinal Cord Pool	2.4
Kidney Pool	0.0	Adrenal Gland	3.2
Fetal Kidney	0.7	Pituitary gland Pool	0.6
Renal ca. 786-0	5.3	Salivary Gland	2.5
Renal ca. A498	4.0	Thyroid (female)	25.3
Renal ca. ACHN	3.0	Pancreatic ca. CAPAN2	5.7
Renal ca. UO-31	15.2	Pancreas Pool	3.0

Table ANF. HASS Panel v1.0

Tissue Name	Rel. Exp.(%) Ag2052, Run 247736616	Rel. Exp.(% Ag2052, Run 248455625	Tissue Name	Rel. Exp.(%) Ag2052, Run 247736616	Rel. Exp.(%) Ag2052, Run 248455625
MCF-7 C1	12.6	7.1	U87-MG F1 (B)	40.3	22.4
MCF-7 C2	12.7	8.6	U87-MG F2	11.1	6.7
MCF-7 C3	10.2	5.6	U87-MG F3	12.2	8.0
MCF-7 C4	16.2	19.5	U87-MG F4	27.0	17.8
MCF-7 C5	13.2	11.0	U87-MG F5	59.0	38.2
MCF-7 C6	13.2	14.6	U87-MG F6	61.1	44.4
MCF-7 C7	12.7	10.4	U87-MG F7	72.7	50.7
MCF-7 C9	9.7	12.9	U87-MG F8	75.3	54.7
MCF-7 C10	15.8	17.1	U87-MG F9	29.9	28.1
MCF-7 C11	2.5	1.8	U87-MG F10	65.1	50.0
MCF-7 C12	9.9	8.0	U87-MG F11	58.2	48.3
MCF-7 C13	12.5	17.1	U87-MG F12	47.0	42.6
MCF-7 C15	5.6	6.5	U87-MG F13	95.3	77.9
MCF-7 C16	14.0	21.5	U87-MG F14	96.6	80.1
MCF-7 C17	10.2	6.9	U87-MG F15	64.6	54.7
T24 D1	25.0	14.4	U87-MG F16	51.8	47.6
T24 D2	33.0	42.0	U87-MG F17	62.0	49.0
T24 D3	29.3	19.1	LnCAP A1	9.4	6.0
T24 D4	39.8	30.6	LnCAP A2	8.1	5.5
T24 D5	28.5	19.5	LnCAP A3	6.3	3.4
T24 D6	32.8	27.2	LnCAP A4	10.4	6.9
T24 D7	18.3	25.9	LnCAP A5	10.0	6.0
T24 D9	12.1	8.5	LnCAP A6	10.0	6.3
T24 D10	23.5	19.2	LnCAP A7	9.2	6.6
T24 D11	13.2	11.7	LnCAP A8	11.5	8.8
T24 D12	24.0	19.2	LnCAP A9	10.8	7.2
T24 D13	8.5	5.8	LnCAP A10	11.0	8.0
T24 D15	10.7	8.0	LnCAP A11	15.7	10.7
T24 D16	6.6	4.7	LnCAP A12	3.5	2.3
T24 D17	12.0	7.4	LnCAP A13	5.7	3.3
CAPaN B1	64.6	52.1	LnCAP A14	3.3	1.7
CAPaN B2	46.3	33.2	LnCAP A15	2.5	1.3
CAPaN B3	13.0	10.7	LnCAP A16	12.5	8.6
CAPaN B4	39.8	30.4	LnCAP A17	12.2	2.5
CAPaN B5	39.5	28.7	Primary Astrocytes	47.3	27.9

CAPaN B6	27.5	25.7	Primary Renal Proximal Tubule Epithelial cell A2	100.0	100.0
CAPaN B7	30.1	31.2	Primary melanocytes A5	40.1	21.8
CAPaN B8	33.2	26.8	126443 - 341 medullo	0.7	0.4
CAPaN B9	38.7	50.0	126444 - 487 medullo	2.2	1.8
CAPaN B10	57.4	51.4	126445 - 425 medullo	1.6	1.0
CAPaN B11	45.1	28.5	126446 - 690 medullo	4.4	2.6
CAPaN B12	31.4	22.7	126447 - 54 adult glioma	33.4	22.2
CAPaN B13	38.7	29.7	126448 - 245 adult glioma	9.4	6.3
CAPaN B14	29.9	22.1	126449 - 317 adult glioma	10.4	6.0
CAPaN B15	32.8	20.7	126450 - 212 glioma	41.5	22.8
CAPaN B16	29.7	16.4	126451 - 456 glioma	17.4	11.3
CAPaN B17	42.3	24.3			

Table ANG. Panel 1.3D

Tissue Name	Rel. Exp.(% Ag2052, Run 166004256	Tissue Name	Rel. Exp.(%) Ag2052, Run 166004256
Liver adenocarcinoma	21.8	Kidney (fetal)	19.2
Pancreas	4.2	Renal ca. 786-0	8.4
Pancreatic ca. CAPAN 2	24.5	Renal ca. A498	26.4
Adrenal gland	11.7	Renal ca. RXF 393	34.4
Thyroid	37.6	Renal ca. ACHN	9.3
Salivary gland	25.3	Renal ca. UO-31	33.7
Pituitary gland	13.8	Renal ca. TK-10	2.8
Brain (fetal)	11.7	Liver	14.0
Brain (whole)	51.4	Liver (fetal)	16.2
Brain (amygdala)	29.5	Liver ca. (hepatoblast) HepG2	33.9
Brain (cerebellum)	24.3	Lung	22.8
Brain (hippocampus)	24.5	Lung (fetal)	10.7
Brain (substantia nigra)	17.8	Lung ca. (small cell) LX-1	25.2
Brain (thalamus)	27.5	Lung ca. (small cell) NCI-H69	2.1
Cerebral Cortex	45.4	Lung ca. (s.cell var.) SHP-77	6.9
Spinal cord	30.4	Lung ca. (large cell)NCI-H460	2.1
glio/astro U87-MG	42.6	Lung ca. (non-sm. cell) A549	4.4
glio/astro U-118-MG	23.5	Lung ca. (non-s.cell) NCI-H23	4.4

astrocytoma SW1783	24.3	Lung ca. (non-s.cell) HOP-62	30.4
neuro*; met SK-N-AS	5.4	Lung ca. (non-s.cl) NCI-H522	3.4
astrocytoma SF-539	43.8	Lung ca. (squam.) SW 900	18.4
astrocytoma SNB-75	21.9	Lung ca. (squam.) NCI-H596	1.9
glioma SNB-19	20.7	Mammary gland	15.5
glioma U251	43.2	Breast ca.* (pl.ef) MCF-7	10.7
glioma SF-295	25.5	Breast ca.* (pl.ef) MDA-MB-231	13.2
Heart (fetal)	15.2	Breast ca.* (pl.ef) T47D	6.0
Heart	13.7	Breast ca. BT-549	100.0
Skeletal muscle (fetal)	8.2	Breast ca. MDA-N	3.7
Skeletal muscle	11.8	Ovary	23.5
Bone marrow	19.5	Ovarian ca. OVCAR-3	14.1
Thymus	7.7	Ovarian ca. OVCAR-4	20.7
Spleen	34.6	Ovarian ca. OVCAR-5	23.5
Lymph node	17.4	Ovarian ca. OVCAR-8	7.8
Colorectal	12.5	Ovarian ca. IGROV-1	5.1
Stomach	8.0	Ovarian ca.* (ascites) SK-OV-3	27.9
Small intestine	12.2	Uterus	11.0
Colon ca. SW480	9.7	Placenta	40.3
Colon ca.* SW620(SW480 met)	5.9	Prostate	8.0
Colon ca. HT29	1.2	Prostate ca.* (bone met)PC-3	8.4
Colon ca. HCT-116	4.8	Testis	4.3
Colon ca. CaCo-2	15.7	Melanoma Hs688(A).T	22.7
Colon ca. tissue(ODO3866)	62.4	Melanoma* (met) Hs688(B).T	21.8
Colon ca. HCC-2998	12.9	Melanoma UACC-62	23.0
Gastric ca.* (liver met) NCI-N87	21.9	Melanoma M14	43.2
Bladder	11.4	Melanoma LOX IMVI	11.2
Trachea	13.1	Melanoma* (met) SK-MEL-5	22.8
Kidney	31.0	Adipose	12.8

Table ANH. Panel 2.2

5

Tissue Name	Rel. Exp. %) Ag2052, Run 174244470	Tissue Name	Rel. Exp.(%) Ag2052, Run 174244470
Normal Colon	3.3	Kidney Margin (OD04348)	13.1
Colon cancer (OD06064)	23.3	Kidney malignant cancer (OD06204B)	1.0
Colon Margin (OD06064)	3.6	Kidney normal adjacent tissue (OD06204E)	9.5
Colon cancer (OD06159)	1.5	Kidney Cancer (OD04450-01)	22.2

Colon Margin (OD06159)	3.6	Kidney Margin (OD04450-03)	4.5
Colon cancer (OD06297-04)	1.3	Kidney Cancer 8120613	0.6
Colon Margin (OD06297-05)	4.7	Kidney Margin 8120614	0.0
CC Gr.2 ascend colon (ODO3921)	1.5	Kidney Cancer 9010320	10.7
CC Margin (ODO3921)	2.6	Kidney Margin 9010321	6.6
Colon cancer metastasis (OD06104)	6.7	Kidney Cancer 8120607	9.7
Lung Margin (OD06104)	6.0	Kidney Margin 8120608	11.4
Colon mets to lung (OD04451-01)	12.8	Normal Uterus	3.1
Lung Margin (OD04451-02)	6.0	Uterine Cancer 064011	3.5
Normal Prostate	2.3	Normal Thyroid	7.2
Prostate Cancer (OD04410)	0.7	Thyroid Cancer 064010	44.8
Prostate Margin (OD04410)	1.2	Thyroid Cancer A302152	**100.0**
Normal Ovary	6.1	Thyroid Margin A302153	7.6
Ovarian cancer (OD06283-03)	4.1	Normal Breast	2.2
Ovarian Margin (OD06283-07)	2.0	Breast Cancer (OD04566)	2.5
Ovarian Cancer 064008	9.2	Breast Cancer 1024	6.3
Ovarian cancer (OD06145)	8.9	Breast Cancer (OD04590-01)	8.5
Ovarian Margin (OD06145)	3.8	Breast Cancer Mets (OD04590-03)	4.4
Ovarian cancer (OD06455-03)	6.1	Breast Cancer Metastasis (OD04655-05)	3.3
Ovarian Margin (OD06455-07)	1.0	Breast Cancer 064006	4.9
Normal Lung	4.9	Breast Cancer 9100266	2.7
Invasive poor diff. lung adeno (ODO4945-01	2.9	Breast Margin 9100265	1.7
Lung Margin (ODO4945-03)	3.2	Breast Cancer A209073	1.5
Lung Malignant Cancer (OD03126)	11.1	Breast Margin A2090734	2.3
Lung Margin (OD03126)	5.1	Breast cancer (OD06083)	4.4
Lung Cancer (OD05014A)	19.6	Breast cancer node metastasis (OD06083)	5.6
Lung Margin (OD05014B)	15.3	Normal Liver	6.9
Lung cancer (OD06081)	3.4	Liver Cancer 1026	8.0
Lung Margin (OD06081)	1.3	Liver Cancer 1025	22.2
Lung Cancer (OD04237-01)	4.6	Liver Cancer 6004-T	13.8
Lung Margin (OD04237-02)	11.1	Liver Tissue 6004-N	4.1
Ocular Melanoma Metastasis	3.5	Liver Cancer 6005-T	21.5
Ocular Melanoma Margin (Liver)	9.8	Liver Tissue 6005-N	51.1
Melanoma Metastasis	5.4	Liver Cancer 064003	13.6
Melanoma Margin (Lung)	5.1	Normal Bladder	2.8
Normal Kidney	3.3	Bladder Cancer 1023	4.8
Kidney Ca, Nuclear grade 2 (OD04338)	5.0	Bladder Cancer A302173	6.1
Kidney Margin (OD04338)	10.6	Normal Stomach	5.3

Kidney Ca Nuclear grade 1/2 (OD04339)	15.0	Gastric Cancer 9060397	6.3
Kidney Margin (OD04339)	11.3	Stomach Margin 9060396	5.0
Kidney Ca, Clear cell type (OD04340)	4.2	Gastric Cancer 9060395	4.6
Kidney Margin (OD04340)	7.2	Stomach Margin 9060394	7.7
Kidney Ca, Nuclear grade 3 (OD04348)	3.1	Gastric Cancer 064005	3.8

Table ANI. Panel 4.1D

5

Tissue Name	Rel. Exp.(% Ag5278, Run 230472911	Tissue Name	Rel. Exp.(%) Ag5278, Run 230472911
Secondary Th1 act	3.4	HUVEC IL-1beta	13.0
Secondary Th2 act	3.3	HUVEC IFN gamma	9.0
Secondary Tr1 act	1.2	HUVEC TNF alpha + IFN gamma	7.4
Secondary Th1 rest	0.4	HUVEC TNF alpha + IL4	2.1
Secondary Th2 rest	0.0	HUVEC IL-11	3.6
Secondary Tr1 rest	0.0	Lung Microvascular EC none	27.7
Primary Th1 act	0.0	Lung Microvascular EC TNFalpha + IL-1beta	8.2
Primary Th2 act	1.1	Microvascular Dermal EC none	4.2
Primary Tr1 act	1.4	Microsvasular Dermal EC TNFalpha + IL-1beta	3.0
Primary Th1 rest	0.5	Bronchial epithelium TNFalpha + IL1beta	9.1
Primary Th2 rest	0.5	Small airway epithelium none	22.1
Primary Tr1 rest	0.9	Small airway epithelium TNFalpha + IL-1beta	33.9
CD45RA CD4 lymphocyte act	5.0	Coronery artery SMC rest	6.2
CD45RO CD4 lymphocyte act	1.6	Coronery artery SMC TNFalpha + IL-1beta	11.3
CD8 lymphocyte act	0.4	Astrocytes rest	2.3
Secondary CD8 lymphocyte rest	1.3	Astrocytes TNFalpha + IL-1beta	3.1
Secondary CD8 lymphocyte act	0.0	KU-812 (Basophil) rest	1.9
CD4 lymphocyte none	0.0	KU-812 (Basophil) PMA/ionomycin	10.9
2ry Th1/Th2/Tr1_anti-CD95 CH11	0.0	CCD1106 (Keratinocytes) none	5.8
LAK cells rest	18.6	CCD1106 (Keratinocytes) TNFalpha + IL-1beta	4.8
LAK cells IL-2	0.6	Liver cirrhosis	1.9

LAK cells IL-2+IL-12	0.0	NCI-H292 none	7.1
LAK cells IL-2+IFN gamma	0.7	NCI-H292 IL-4	8.4
LAK cells IL-2+ IL-18	0.9	NCI-H292 IL-9	7.0
LAK cells PMA/ionomycin	62.4	NCI-H292 IL-13	5.6
NK Cells IL-2 rest	1.0	NCI-H292 IFN gamma	3.6
Two Way MLR 3 day	9.4	HPAEC none	9.1
Two Way MLR 5 day	3.9	HPAEC TNF alpha + IL-1 beta	28.3
Two Way MLR 7 day	2.3	Lung fibroblast none	9.3
PBMC rest	0.6	Lung fibroblast TNF alpha + IL-1 beta	12.2
PBMC PWM	1.1	Lung fibroblast IL-4	3.9
PBMC PHA-L	2.2	Lung fibroblast IL-9	11.8
Ramos (B cell) none	0.0	Lung fibroblast IL-13	5.4
Ramos (B cell) ionomycin	0.0	Lung fibroblast IFN gamma	19.5
B lymphocytes PWM	0.0	Dermal fibroblast CCD1070 rest	32.1
B lymphocytes CD40L and IL-4	1.4	Dermal fibroblast CCD1070 TNF alpha	66.0
EOL-1 dbcAMP	1.4	Dermal fibroblast CCD1070 IL-1 beta	21.8
EOL-1 dbcAMP PMA/ionomycin	1.4	Dermal fibroblast IFN gamma	42.3
Dendritic cells none	**100.0**	Dermal fibroblast IL-4	45.1
Dendritic cells LPS	34.9	Dermal Fibroblasts rest	15.7
Dendritic cells anti-CD40	44.8	Neutrophils TNFa+LPS	0.0
Monocytes rest	1.4	Neutrophils rest	0.6
Monocytes LPS	19.9	Colon	0.0
Macrophages rest	12.5	Lung	1.4
Macrophages LPS	11.2	Thymus	0.0
HUVEC none	5.9	Kidney	12.8
HUVEC starved	11.7		

Table ANJ. Panel 4D

5

Tissue Name	Rel. Exp.() Ag2052, Run 161706487	Tissue Name	Rel. Exp.(%) Ag2052, Run 161706487
Secondary Th1 act	2.6	HUVEC IL-1beta	2.1
Secondary Th2 act	1.7	HUVEC IFN gamma	5.2
Secondary Tr1 act	1.9	HUVEC TNF alpha + IFN gamma	5.7
Secondary Th1 rest	0.3	HUVEC TNF alpha + IL4	4.5
Secondary Th2 rest	0.5	HUVEC IL-11	2.6
Secondary Tr1 rest	0.6	Lung Microvascular EC none	9.9

Primary Th1 act	1.4	Lung Microvascular EC TNFalpha + IL-1beta	10.0
Primary Th2 act	0.7	Microvascular Dermal EC none	16.6
Primary Tr1 act	1.2	Microsvasular Dermal EC TNFalpha + IL-1beta	9.2
Primary Th1 rest	2.2	Bronchial epithelium TNFalpha + IL1beta	3.1
Primary Th2 rest	1.4	Small airway epithelium none	12.5
Primary Tr1 rest	0.2	Small airway epithelium TNFalpha + IL-1beta	46.0
CD45RA CD4 lymphocyte act	4.2	Coronery artery SMC rest	5.4
CD45RO CD4 lymphocyte act	1.4	Coronery artery SMC TNFalpha + IL-1beta	4.3
CD8 lymphocyte act	0.3	Astrocytes rest	2.2
Secondary CD8 lymphocyte rest	1.4	Astrocytes TNFalpha + IL-1beta	2.0
Secondary CD8 lymphocyte act	0.4	KU-812 (Basophil) rest	1.5
CD4 lymphocyte none	0.4	KU-812 (Basophil) PMA/ionomycin	11.0
2ry Th1/Th2/Tr1_anti-CD95 CH11	0.8	CCD1106 (Keratinocytes) none	3.1
LAK cells rest	43.2	CCD1106 (Keratinocytes) TNFalpha + IL-1beta	0.8
LAK cells IL-2	0.8	Liver cirrhosis	1.5
LAK cells IL-2+IL-12	1.8	Lupus kidney	0.7
LAK cells IL-2+IFN gamma	3.2	NCI-H292 none	5.8
LAK cells IL-2+ IL-18	2.1	NCI-H292 IL-4	5.5
LAK cells PMA/ionomycin	26.2	NCI-H292 IL-9	7.4
NK Cells IL-2 rest	0.3	NCI-H292 IL-13	2.7
Two Way MLR 3 day	9.2	NCI-H292 IFN gamma	3.3
Two Way MLR 5 day	9.3	HPAEC none	5.6
Two Way MLR 7 day	2.0	HPAEC TNF alpha + IL-1 beta	10.7
PBMC rest	1.0	Lung fibroblast none	6.3
PBMC PWM	5.3	Lung fibroblast TNF alpha + IL-1 beta	6.3
PBMC PHA-L	5.0	Lung fibroblast IL-4	10.4
Ramos (B cell) none	0.0	Lung fibroblast IL-9	8.1
Ramos (B cell) ionomycin	0.0	Lung fibroblast IL-13	5.6
B lymphocytes PWM	2.2	Lung fibroblast IFN gamma	15.4
B lymphocytes CD40L and IL-4	1.2	Dermal fibroblast CCD1070 rest	15.5
EOL-1 dbcAMP	0.7	Dermal fibroblast CCD1070 TNF alpha	18.9
EOL-1 dbcAMP PMA/ionomycin	1.5	Dermal fibroblast CCD1070 IL-1 beta	11.1
Dendritic cells none	66.9	Dermal fibroblast IFN gamma	19.6
Dendritic cells LPS	37.6	Dermal fibroblast IL-4	21.2
Dendritic cells anti-CD40	77.9	IBD Colitis 2	0.2

Monocytes rest	5.1	IBD Crohn's	0.5
Monocytes LPS	17.2	Colon	3.9
Macrophages rest	**100.0**	Lung	19.8
Macrophages LPS	40.9	Thymus	12.9
HUVEC none	5.3	Kidney	2.4
HUVEC starved	10.6		

Table ANK. Panel 5 Islet

5

Tissue Name	Rel. Exp.(%) Ag2052, un 279370795	Tissue Name	Rel. Exp.(%) Ag2052, Run 279370795
97457_Patient-02go_adipose	15.6	94709_Donor 2 AM - A_adipose	24.7
97476_Patient-07sk_skeletal muscle	0.0	94710_Donor 2 AM - B_adipose	24.7
97477_Patient-07ut_uterus	22.1	94711_Donor 2 AM - C_adipose	14.7
97478_Patient-07pl_placenta	13.1	94712_Donor 2 AD - A_adipose	64.2
99167_Bayer Patient 1	17.6	94713_Donor 2 AD - B_adipose	89.5
97482_Patient-08ut_uterus	15.3	94714_Donor 2 AD - C_adipose	66.4
97483_Patient-08pl_placenta	11.6	94742_Donor 3 U - A_Mesenchymal Stem Cells	17.3
97486_Patient-09sk_skeletal muscle	4.8	94743_Donor 3 U - B_Mesenchymal Stem Cells	23.2
97487_Patient-09ut_uterus	15.5	94730_Donor 3 AM - A_adipose	54.0
97488_Patient-09pl_placenta	7.9	94731_Donor 3 AM - B_adipose	76.3
97492_Patient-10ut_uterus	14.5	94732_Donor 3 AM - C_adipose	59.9
97493_Patient-10pl_placenta	23.8	94733_Donor 3 AD - A_adipose	**100.0**
97495_Patient-11go_adipose	11.9	94734_Donor 3 AD - B_adipose	92.0
97496_Patient-11sk_skeletal muscle	3.2	94735_Donor 3 AD - C_adipose	32.1
97497_Patient-11ut_uterus	36.9	77138_Liver_HepG2untreated	62.9
97498_Patient-11pl_placenta	7.0	73556_Heart_Cardiac stromal cells (primary)	0.3
97500_Patient-12go_adipose	17.2	81735_Small Intestine	10.9
97501_Patient-12sk_skeletal muscle	8.4	72409_Kidney_Proximal Convoluted Tubule	23.7
97502_Patient-12ut_uterus	25.2	82685_Small intestine_Duodenum	9.3
97503_Patient-12pl_placenta	23.8	90650_Adrenal_Adrenocortical adenoma	8.4
94721_Donor 2 U - A_Mesenchymal Stem Cells	61.6	72410_Kidney_HRCE	40.1
94722_Donor 2 U - B_Mesenchymal Stem Cells	45.1	72411_Kidney_HRE	13.5

94723_Donor 2 U - C_Mesenchymal Stem Cells	53.2	73139_Uterus_Uterine smooth muscle cells	61.1

AI_comprehensive panel_v1.0 Summary: Ag2052 Highest expression of this gene is detected in synovium from an orthoarthritis (OA) patient (CT=20.3). High levels of expression of this gene are detected in samples derived from normal and orthoarthitis/ rheumatoid arthritis bone and adjacent bone, cartilage, synovium and synovial fluid samples, from normal lung, COPD lung, emphysema, atopic asthma, asthma, allergy, Crohn's disease (normal matched control and diseased), ulcerative colitis(normal matched control and diseased), and psoriasis (normal matched control and diseased). Therefore, therapeutic modulation of this gene product may ameliorate symptoms/conditions associated with autoimmune and inflammatory disorders including psoriasis, allergy, asthma, inflammatory bowel disease, rheumatoid arthritis and osteoarthritis.

CNS_neurodegeneration_v1.0 Summary: Ag5277/Ag5278 Expression of this gene is low/undetectable (CTs > 35) across all of the samples on this panel.

General_screening_panel_v1.5 Summary: Ag5278 Highest expression of this gene is detected in breast cancer BT-549 cell line (CT=29). Moderate levels of expression of this gene is also seen in cluster of cancer cell lines derived from pancreatic, gastric, colon, lung, liver, renal, breast, ovarian, melanoma and brain cancers. In addition, moderate to low levels of expression of this gene is also seen in all the regions of brain, in tissues with metabolic/endocrine functions such as pancreas, adrenal gland, thyroid, fetal liver and colon. Please see panel 1.3D for further discussion of this gene.

Ag5277 Expression of this gene is low/undetectable (CTs > 35) across all of the samples on this panel.

HASS Panel v1.0 Summary: Ag2052 Two experiments with same probe and primer sets are in excellent agreement. This gene shows wide spread expression in this panel, with highest expression in primary renal proximal tubular epithelial cells cultured in vitro (CTs=20-22). The expression of this gene is also higher in the glioblastoma type of brain cancer compared to the medulloblastoma suggesting that it may play a role in glioblastoma development than medulloblastomas. Expression is also induced in the U87-MG(cells when they are deprived of nutrients, oxygen and exposed to an acidic pH than in the control population (comparing the control U87-MG F4 with U87-MG F5, F7, F10). This suggests that the serum-starved, hypoxic and acidotic regions of brain cancers

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may express this gene at a higher level and that this may be used as a marker for these 3 7 3 regions.

Panel 1.3D Summary: Ag2052 This gene shows a widespread expression in this panel. Highest expression of this gene is detected in breast cancer BT-549 cell line

5   (CT=24.9). High levels of expression of this gene is also seen in cluster of cancer cell lines derived from pancreatic, gastric, colon, lung, liver, renal, breast, ovarian, prostate, melanoma and brain cancers. Thus, expression of this gene could be used as a marker to detect the presence of these cancers. Furthermore, therapeutic modulation of the expression or function of this gene may be effective in the treatment of pancreatic, gastric, colon, lung,

10   liver, renal, breast, ovarian, prostate, melanoma and brain cancers.

Among tissues with metabolic or endocrine function, this gene is expressed at high levels in pancreas, adipose, adrenal gland, thyroid, pituitary gland, skeletal muscle, heart, liver and the gastrointestinal tract. Therefore, therapeutic modulation of the activity of this gene may prove useful in the treatment of endocrine/metabolically related diseases, such as

15   obesity and diabetes.

In addition, this gene is expressed at high levels in all regions of the central nervous system examined, including amygdala, hippocampus, substantia nigra, thalamus, cerebellum, cerebral cortex, and spinal cord. Therefore, therapeutic modulation of this gene product may be useful in the treatment of central nervous system disorders such as

20   Alzheimer's disease, Parkinson's disease, epilepsy, multiple sclerosis, schizophrenia and depression.

Panel 2.2 Summary: Ag2052 Highest expression of this gene is detected in thyroid cancer (CT=23.9). High to moderate levels of expression of this gene is also seen in normal and cancer samples derived from melanoma, colon, gastric, bladder, liver, breast,

25   thyroid, uterine, kidney, lung, ovarian and prostate cancers. Interestingly, higher levels of expression of this gene is associated with kidney and thyroid cancers as compared to corresponding normal tissue. Therefore, expression of this gene may bay used as diagnostic marker to detect the presence of these cancers. Furthermore, therapeutic modulation of this gene may be useful in the treatment of melanoma, colon, gastric, bladder, liver, breast,

30   thyroid, uterine, kidney, lung, ovarian and prostate cancers.

Panel 4.1D Summary: Ag5278 Highest levels of expression of this gene is detected in resting dendritic cells (CT=32). Moderate to low levels of expression of this gene is also seen in activated dendrict cells, PMA/ionomycin stimulated LAK cells, LPS

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activated macrophage, lung microvascular endothelial cells, activated HPAEC cells, small
airway epithelium, and dermal fibroblasts. Therefore, therapeutic modulation of this gene
or its protein product may alter the functions associated with these cell types and would be
beneficial in the treatment of autoimmune and inflammatory diseases such as asthma,

5   allergies, inflammatory bowel disease, lupus erythematosus, psoriasis, rheumatoid arthritis,
and osteoarthritis.

Ag5277 Expression of this gene is low/undetectable (CTs > 35) across all of the
samples on this panel.

Panel 4D Summary: Ag2052 Highest expression of this gene is detected in resting
10   macrophage (CT=21). This gene is expressed at high to moderate levels in a wide range of
cell types of significance in the immune response in health and disease. These cells include
members of the T-cell, B-cell, dendritic cells, endothelial cell, macrophage/monocyte, and
peripheral blood mononuclear cell family, as well as epithelial and fibroblast cell types
from lung and skin, and normal tissues represented by colon, lung, thymus and kidney. This
15   ubiquitous pattern of expression suggests that this gene product may be involved in
homeostatic processes for these and other cell types and tissues. This pattern is in
agreement with the expression profile in General_screening_panel_v1.3 and also suggests a
role for the gene product in cell survival and proliferation. Therefore, modulation of the
gene product with a functional therapeutic may lead to the alteration of functions associated
20   with these cell types and lead to improvement of the symptoms of patients suffering from
autoimmune and inflammatory diseases such as asthma, allergies, inflammatory bowel
disease, lupus erythematosus, psoriasis, rheumatoid arthritis, and osteoarthritis.

Panel 5 Islet Summary: Ag2052 Highest expression of this gene is detected in a
differentiated adipose tissue (CT=24.4). Moderate to high levels of expression is seen in
25   placenta, uterus, adipose, skeletal muscle, small intestine, heart and kidney. This gene
shows a ubiquitous expression which correlates to the expression in panel 1.3D. Please see
panel 1.3D for further discussion of this gene.

AO. CG56836-04: Cathepsin B.

Expression of gene CG56836-04 was assessed using the primer-probe set Ag5264,
30   described in Table AOA. Results of the RTQ-PCR runs are shown in Tables AOB, AOC
and AOD.

Table AOA. Probe Name Ag5264

Primers		Length	Start Position	SEQ ID No
Forward	5'-tcctgctgggtttctggt-3'	18	455	411
Probe	TET-5'-ccgtactccatccctccctgtgagc-3'-TAMRA	25	503	412
Reverse	5'-tgtttgtaggtcgggctgta-3'	20	605	413

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Table AOB. CNS_neurodegeneration_v1.0

Tissue Name	Rel. Exp.(%) Ag5264, Run 230512807	issue Name	Rel. Exp.(%) Ag5264, Run 230512807
AD 1 Hippo	10.2	Control (Path) 3 Temporal Ctx	3.6
AD 2 Hippo	32.5	Control (Path) 4 Temporal Ctx	18.4
AD 3 Hippo	9.3	AD 1 Occipital Ctx	14.7
AD 4 Hippo	3.8	AD 2 Occipital Ctx (Missing)	0.0
AD 5 hippo	94.0	AD 3 Occipital Ctx	7.3
AD 6 Hippo	66.9	AD 4 Occipital Ctx	13.4
Control 2 Hippo	25.0	AD 5 Occipital Ctx	15.3
Control 4 Hippo	13.0	AD 6 Occipital Ctx	39.0
Control (Path) 3 Hippo	4.0	Control 1 Occipital Ctx	5.9
AD 1 Temporal Ctx	9.8	Control 2 Occipital Ctx	53.6
AD 2 Temporal Ctx	25.2	Control 3 Occipital Ctx	8.4
AD 3 Temporal Ctx	3.9	Control 4 Occipital Ctx	6.3
AD 4 Temporal Ctx	7.5	Control (Path) 1 Occipital Ctx	83.5
AD 5 Inf Temporal Ctx	74.7	Control (Path) 2 Occipital Ctx	6.0
AD 5 SupTemporal Ctx	43.8	Control (Path) 3 Occipital Ctx	1.7
AD 6 Inf Temporal Ctx	71.2	Control (Path) 4 Occipital Ctx	13.1
AD 6 Sup Temporal Ctx	41.8	Control 1 Parietal Ctx	2.9
Control 1 Temporal Ctx	5.9	Control 2 Parietal Ctx	30.1
Control 2 Temporal Ctx	45.1	Control 3 Parietal Ctx	12.3
Control 3 Temporal Ctx	12.0	Control (Path) 1 Parietal Ctx	100.0
Control 4 Temporal Ctx	6.7	Control (Path) 2 Parietal Ctx	12.6
Control (Path) 1 Temporal Ctx	47.3	Control (Path) 3 Parietal Ctx	2.5
Control (Path) 2 Temporal Ctx	15.9	Control (Path) 4 Parietal Ctx	44.1

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Table AOC. General screening panel v1.5

Tissue Name	Rel. Exp.(%) Ag5264, Run 232936651	issue Name	Rel. Exp.(%) Ag5264, Run 232936651
Adipose	0.7	Renal ca. TK-10	3.6
Melanoma* Hs688(A).T	19.5	Bladder	3.8
Melanoma* Hs688(B).T	9.0	Gastric ca. (liver met.) NCI-N87	10.2
Melanoma* M14	24.7	Gastric ca. KATO III	5.5
Melanoma* LOXIMVI	15.6	Colon ca. SW-948	1.2
Melanoma* SK-MEL-5	9.7	Colon ca. SW480	7.0
Squamous cell carcinoma SCC-4	3.1	Colon ca.* (SW480 met) SW620	2.0
Testis Pool	0.4	Colon ca. HT29	0.6
Prostate ca.* (bone met) PC-3	2.0	Colon ca. HCT-116	3.1
Prostate Pool	0.6	Colon ca. CaCo-2	5.2
Placenta	3.7	Colon cancer tissue	8.6
Uterus Pool	0.2	Colon ca. SW1116	2.4
Ovarian ca. OVCAR-3	6.7	Colon ca. Colo-205	4.1
Ovarian ca. SK-OV-3	7.2	Colon ca. SW-48	1.3
Ovarian ca. OVCAR-4	4.2	Colon Pool	1.2
Ovarian ca. OVCAR-5	6.2	Small Intestine Pool	0.7
Ovarian ca. IGROV-1	1.5	Stomach Pool	1.3
Ovarian ca. OVCAR-8	2.2	Bone Marrow Pool	0.7
Ovary	1.4	Fetal Heart	0.5
Breast ca. MCF-7	2.7	Heart Pool	1.3
Breast ca. MDA-MB-231	4.9	Lymph Node Pool	2.2
Breast ca. BT 549	100.0	Fetal Skeletal Muscle	0.3
Breast ca. T47D	1.3	Skeletal Muscle Pool	1.3
Breast ca. MDA-N	1.1	Spleen Pool	1.2
Breast Pool	1.7	Thymus Pool	0.9
Trachea	3.0	CNS cancer (glio/astro) U87-MG	12.6
Lung	0.2	CNS cancer (glio/astro) U-118-MG	9.0
Fetal Lung	1.6	CNS cancer (neuro;met) SK-N-AS	2.1
Lung ca. NCI-N417	0.2	CNS cancer (astro) SF-539	7.4
Lung ca. LX-1	4.5	CNS cancer (astro) SNB-75	22.5
Lung ca. NCI-H146	0.2	CNS cancer (glio) SNB-19	1.7
Lung ca. SHP-77	1.6	CNS cancer (glio) SF-295	15.6
Lung ca. A549	4.1	Brain (Amygdala) Pool	1.4
Lung ca. NCI-H526	0.2	Brain (cerebellum)	5.6
Lung ca. NCI-H23	2.2	Brain (fetal)	1.0
Lung ca. NCI-H460	1.2	Brain (Hippocampus) Pool	1.3
Lung ca. HOP-62	5.6	Cerebral Cortex Pool	1.6

Lung ca. NCI-H522	1.4	Brain (Substantia nigra) Pool	1.5
Liver	1.7	Brain (Thalamus) Pool	2.1
Fetal Liver	4.9	Brain (whole)	3.1
Liver ca. HepG2	4.9	Spinal Cord Pool	1.6
Kidney Pool	2.4	Adrenal Gland	2.1
Fetal Kidney	1.0	Pituitary gland Pool	0.4
Renal ca. 786-0	1.0	Salivary Gland	1.6
Renal ca. A498	1.7	Thyroid (female)	16.7
Renal ca. ACHN	4.0	Pancreatic ca. CAPAN2	5.6
Renal ca. UO-31	11.2	Pancreas Pool	2.8

Table AOD. Panel 4.1D

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Tissue Name	Rel. Exp.(% Ag5264, Run 230472870	Tissue Name	Rel. Exp.(%) Ag5264, Run 230472870
Secondary Th1 act	4.0	HUVEC IL-1beta	9.2
Secondary Th2 act	3.3	HUVEC IFN gamma	7.2
Secondary Tr1 act	1.2	HUVEC TNF alpha + IFN gamma	4.6
Secondary Th1 rest	0.3	HUVEC TNF alpha + IL4	5.1
Secondary Th2 rest	0.2	HUVEC IL-11	4.5
Secondary Tr1 rest	0.2	Lung Microvascular EC none	32.5
Primary Th1 act	0.5	Lung Microvascular EC TNFalpha + IL-1beta	10.3
Primary Th2 act	0.7	Microvascular Dermal EC none	4.2
Primary Tr1 act	1.0	Microsvasular Dermal EC TNFalpha + IL-1beta	2.8
Primary Th1 rest	0.2	Bronchial epithelium TNFalpha + IL1beta	11.5
Primary Th2 rest	0.3	Small airway epithelium none	15.8
Primary Tr1 rest	0.2	Small airway epithelium TNFalpha + IL-1beta	20.2
CD45RA CD4 lymphocyte act	4.6	Coronery artery SMC rest	6.0
CD45RO CD4 lymphocyte act	1.7	Coronery artery SMC TNFalpha + IL-1beta	5.1
CD8 lymphocyte act	0.3	Astrocytes rest	1.5
Secondary CD8 lymphocyte rest	1.1	Astrocytes TNFalpha + IL-1beta	1.9
Secondary CD8 lymphocyte act	0.3	KU-812 (Basophil) rest	1.7
CD4 lymphocyte none	0.1	KU-812 (Basophil) PMA/ionomycin	8.9
2ry Th1/Th2/Tr1_anti-CD95 CH11	0.8	CCD1106 (Keratinocytes) none	6.8

LAK cells rest	39.2	CCD1106 (Keratinocytes) TNFalpha + IL-1beta	5.0
LAK cells IL-2	0.6	Liver cirrhosis	3.8
LAK cells IL-2+IL-12	0.1	NCI-H292 none	3.6
LAK cells IL-2+IFN gamma	0.3	NCI-H292 IL-4	4.7
LAK cells IL-2+ IL-18	0.3	NCI-H292 IL-9	5.4
LAK cells PMA/ionomycin	54.3	NCI-H292 IL-13	3.3
NK Cells IL-2 rest	0.6	NCI-H292 IFN gamma	2.4
Two Way MLR 3 day	9.0	HPAEC none	3.7
Two Way MLR 5 day	3.4	HPAEC TNF alpha + IL-1 beta	27.0
Two Way MLR 7 day	1.3	Lung fibroblast none	10.7
PBMC rest	0.4	Lung fibroblast TNF alpha + IL-1 beta	10.4
PBMC PWM	0.7	Lung fibroblast IL-4	4.5
PBMC PHA-L	2.7	Lung fibroblast IL-9	8.2
Ramos (B cell) none	0.0	Lung fibroblast IL-13	2.2
Ramos (B cell) ionomycin	0.0	Lung fibroblast IFN gamma	16.0
B lymphocytes PWM	0.5	Dermal fibroblast CCD1070 rest	17.6
B lymphocytes CD40L and IL-4	1.3	Dermal fibroblast CCD1070 TNF alpha	16.6
EOL-1 dbcAMP	1.0	Dermal fibroblast CCD1070 IL-1 beta	16.7
EOL-1 dbcAMP PMA/ionomycin	0.9	Dermal fibroblast IFN gamma	31.6
Dendritic cells none	100.0	Dermal fibroblast IL-4	20.3
Dendritic cells LPS	31.9	Dermal Fibroblasts rest	14.6
Dendritic cells anti-CD40	36.3	Neutrophils TNFa+LPS	0.2
Monocytes rest	1.4	Neutrophils rest	0.2
Monocytes LPS	40.9	Colon	0.0
Macrophages rest	26.1	Lung	1.4
Macrophages LPS	16.7	Thymus	0.2
HUVEC none	4.7	Kidney	9.7
HUVEC starved	5.8		

CNS_neurodegeneration_v1.0 Summary: Ag5264 This panel confirms the expression of this gene at low levels in the brains of an independent group of individuals. However, no differential expression of this gene was detected between Alzheimer's diseased postmortem brains and those of non-demented controls in this experiment. Please see Panel 1.5 for a discussion of the potential utility of this gene in treatment of central nervous system disorders.

General_screening_panel_v1.5 Summary: Ag5264 Highest expression of this gene is detected in breast cancer BT-549 cell line (CT=25). Moderate levels of expression

of this gene is also seen in cluster of cancer cell lines derived from pancreatic, gastric, colon, lung, liver, renal, breast, ovarian, prostate, melanoma and brain cancers. Thus, expression of this gene could be used as a marker to detect the presence of these cancers. Furthermore, therapeutic modulation of the expression or function of this gene may be

5   effective in the treatment of pancreatic, gastric, colon, lung, liver, renal, breast, ovarian, prostate, melanoma and brain cancers.

Among tissues with metabolic or endocrine function, this gene is expressed at moderate levels in pancreas, adipose, adrenal gland, thyroid, pituitary gland, skeletal muscle, heart, liver and the gastrointestinal tract. Therefore, therapeutic modulation of the

10   activity of this gene may prove useful in the treatment of endocrine/metabolically related diseases, such as obesity and diabetes.

In addition, this gene is expressed at moderate levels in all regions of the central nervous system examined, including amygdala, hippocampus, substantia nigra, thalamus, cerebellum, cerebral cortex, and spinal cord. Therefore, therapeutic modulation of this gene

15   product may be useful in the treatment of central nervous system disorders such as Alzheimer's disease, Parkinson's disease, epilepsy, multiple sclerosis, schizophrenia and depression.

Panel 4.1D Summary: Ag5264 Highest levels of expression of this gene is detected in resting dendritic cells (CT=28.7). Moderate to low levels of expression of this

20   gene is also seen in activated dendritic cells, resting and PMA/ionomycin stimulated LAK cells, monocytes, macrophage, different types of endothelial cells, small airway epithelium, lung and dermal fibroblasts and normal tissue represent by lung and kidney. This gene is upregulated in LPS treated monocytes, cytokine treated HPAEC, and activated secondary Th1, Th2 cells. Therefore, therapeutic modulation of this gene or its protein product may

25   alter the functions associated with these cell types and would be beneficial in the treatment of autoimmune and inflammatory diseases such as asthma, allergies, inflammatory bowel disease, lupus erythematosus, psoriasis, rheumatoid arthritis, and osteoarthritis.

AP. CG57284-03: RAS-RELATED PROTEIN RAB-5C.

Expression of gene CG57284-03 was assessed using the primer-probe set Ag6892,

30   described in Table APA. Results of the RTQ-PCR runs are shown in Tables APB and APC. Please note that this sequence represents a full-length physical clone.

Table APA. Probe Name Ag6892

Primers		Length	Start Position	SEQ ID No
Forward	5'-gtgtcatccaggcagacagtct-3'	22	473	414
Probe	TET-5'-ccgctccaattgtgctctcctggtact-3'-TAMRA	27	507	415
Reverse	5'-cgctttgtcaagggacagttt-3'	21	538	416

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Table APB. General screening panel v1.6

Tissue Name	Rel. Exp.(%) Ag6892, Run 278388295	Tissue Name	Rel. Exp.(%) Ag6892, Run 278388295
Adipose	11.0	Renal ca. TK-10	41.5
Melanoma* Hs688(A).T	37.4	Bladder	19.1
Melanoma* Hs688(B).T	33.0	Gastric ca. (liver met.) NCI-N87	26.4
Melanoma* M14	85.3	Gastric ca. KATO III	93.3
Melanoma* LOXIMVI	48.6	Colon ca. SW-948	15.7
Melanoma* SK-MEL-5	49.7	Colon ca. SW480	62.4
Squamous cell carcinoma SCC-4	28.5	Colon ca.* (SW480 met) SW620	9.5
Testis Pool	10.1	Colon ca. HT29	20.7
Prostate ca.* (bone met) PC-3	0.0	Colon ca. HCT-116	48.0
Prostate Pool	10.6	Colon ca. CaCo-2	49.7
Placenta	22.4	Colon cancer tissue	19.3
Uterus Pool	4.8	Colon ca. SW1116	6.7
Ovarian ca. OVCAR-3	18.9	Colon ca. Colo-205	13.3
Ovarian ca. SK-OV-3	63.3	Colon ca. SW-48	16.5
Ovarian ca. OVCAR-4	17.4	Colon Pool	15.5
Ovarian ca. OVCAR-5	41.5	Small Intestine Pool	8.7
Ovarian ca. IGROV-1	18.4	Stomach Pool	8.0
Ovarian ca. OVCAR-8	13.8	Bone Marrow Pool	8.5
Ovary	10.6	Fetal Heart	5.9
Breast ca. MCF-7	33.2	Heart Pool	6.3
Breast ca. MDA-MB-231	46.0	Lymph Node Pool	16.4
Breast ca. BT 549	37.4	Fetal Skeletal Muscle	5.4
Breast ca. T47D	35.1	Skeletal Muscle Pool	1.6
Breast ca. MDA-N	22.2	Spleen Pool	8.8
Breast Pool	12.7	Thymus Pool	8.7
Trachea	12.0	CNS cancer (glio/astro) U87-MG	35.4
Lung	2.5	CNS cancer (glio/astro) U-118-MG	55.9

Fetal Lung	32.5	CNS cancer (neuro, met) SK-N-AS	52.1
Lung ca. NCI-N417	5.4	CNS cancer (astro) SF-539	28.9
Lung ca. LX-1	20.2	CNS cancer (astro) SNB-75	52.9
Lung ca. NCI-H146	8.6	CNS cancer (glio) SNB-19	21.2
Lung ca. SHP-77	20.2	CNS cancer (glio) SF-295	100.0
Lung ca. A549	51.1	Brain (Amygdala) Pool	10.6
Lung ca. NCI-H526	5.6	Brain (cerebellum)	49.0
Lung ca. NCI-H23	23.7	Brain (fetal)	25.9
Lung ca. NCI-H460	19.1	Brain (Hippocampus) Pool	13.0
Lung ca. HOP-62	21.0	Cerebral Cortex Pool	17.3
Lung ca. NCI-H522	31.4	Brain (Substantia nigra) Pool	11.2
Liver	5.7	Brain (Thalamus) Pool	19.6
Fetal Liver	19.8	Brain (whole)	23.0
Liver ca. HepG2	10.3	Spinal Cord Pool	12.5
Kidney Pool	15.9	Adrenal Gland	24.8
Fetal Kidney	14.0	Pituitary gland Pool	2.7
Renal ca. 786-0	24.3	Salivary Gland	11.3
Renal ca. A498	21.9	Thyroid (female)	9.8
Renal ca. ACHN	22.2	Pancreatic ca. CAPAN2	24.8
Renal ca. UO-31	35.4	Pancreas Pool	8.1

Table APC. Panel 5 Islet

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Tissue Name	Rel. Exp.(%) Ag6892, Run 305424859	Tissue Name	Rel. Exp.(%) Ag6892, Run 305424859
97457_Patient-02go_adipose	4.5	94709_Donor 2 AM - A_adipose	44.1
97476_Patient-07sk_skeletal muscle	0.0	94710_Donor 2 AM - B_adipose	30.8
97477_Patient-07ut_uterus	8.2	94711_Donor 2 AM - C_adipose	21.0
97478_Patient-07pl_placenta	13.1	94712_Donor 2 AD - A_adipose	48.0
99167_Bayer Patient 1	23.2	94713_Donor 2 AD - B_adipose	54.0
97482_Patient-08ut_uterus	7.7	94714_Donor 2 AD - C_adipose	50.3
97483_Patient-08pl_placenta	18.9	94742_Donor 3 U - A_Mesenchymal Stem Cells	14.7
97486_Patient-09sk_skeletal muscle	4.4	94743_Donor 3 U - B_Mesenchymal Stem Cells	10.4
97487_Patient-09ut_uterus	19.6	94730_Donor 3 AM - A_adipose	53.2
97488_Patient-09pl_placenta	11.3	94731_Donor 3 AM - B_adipose	74.2
97492_Patient-10ut_uterus	12.2	94732_Donor 3 AM - C_adipose	58.6
97493_Patient-10pl_placenta	34.9	94733_Donor 3 AD - A_adipose	64.6
97495_Patient-11go_adipose	9.2	94734_Donor 3 AD - B_adipose	100.0

97496_Patient-11sk_skeletal muscle	3.8	94735_Donor 3 AD - C_adipose	20.4
97497_Patient-11ut_uterus	25.0	77138_Liver_HepG2untreated	71.2
97498_Patient-11pl_placenta	8.8	73556_Heart_Cardiac stromal cells (primary)	18.6
97500_Patient-12go_adipose	10.4	81735_Small Intestine	12.4
97501_Patient-12sk_skeletal muscle	12.7	72409_Kidney_Proximal Convoluted Tubule	81.2
97502_Patient-12ut_uterus	18.9	82685_Small intestine_Duodenum	8.1
97503_Patient-12pl_placenta	17.8	90650_Adrenal_Adrenocortical adenoma	4.8
94721_Donor 2 U - A_Mesenchymal Stem Cells	27.9	72410_Kidney_HRCE	37.9
94722_Donor 2 U - B_Mesenchymal Stem Cells	25.7	72411_Kidney_HRE	18.8
94723_Donor 2 U - C_Mesenchymal Stem Cells	30.4	73139_Uterus_Uterine smooth muscle cells	48.0

General_screening_panel_v1.6 Summary: Ag6892 Highest expression of this gene is seen in a brain cancer cell line (CT=24.1). This gene is ubiquitously expressed in this panel, with high levels of expression seen in brain, colon, gastric, lung, breast, ovarian, and melanoma cancer cell lines. This expression profile suggests a role for this gene product in cell survival and proliferation. Modulation of this gene product may be useful in the treatment of cancer.

Among tissues with metabolic function, this gene is expressed at high levels in pituitary, adipose, adrenal gland, pancreas, thyroid, and adult and fetal skeletal muscle, heart, and liver. This widespread expression among these tissues suggests that this gene product may play a role in normal neuroendocrine and metabolic function and that disregulated expression of this gene may contribute to neuroendocrine disorders or metabolic diseases, such as obesity and diabetes.

This gene is also expressed at high levels in the CNS, including the hippocampus, thalamus, substantia nigra, amygdala, cerebellum and cerebral cortex. Therefore, therapeutic modulation of the expression or function of this gene may be useful in the treatment of neurologic disorders, such as Alzheimer's disease, Parkinson's disease, schizophrenia, multiple sclerosis, stroke and epilepsy.

In addition, this gene is expressed at much higher levels in fetal lung tissue (CT=25.7) when compared to expression in the adult counterpart (CT=29.4). Thus,

expression of this gene may be used to differentiate between the fetal and adult source of this tissue.

Panel 5 Islet Summary: Ag6892 Highest expression is seen in adipose (CT=26), with nearly ubiquitous expression seen across the samples on this panel. High to moderate levels of expression are seen in metabolic tissues, including skeletal muscle, adipose, and placenta, in agreement with Panel 1.6. Please see that panel for discussion of this gene in metabolic disease.

AQ. CG57308-02: Sulfonylurea Receptor 1 Splice Variant.

Expression of gene CG57308-02 was assessed using the primer-probe set Ag7558, described in Table AQA. Results of the RTQ-PCR runs are shown in Tables AQB and AQC.

Table AQA. Probe Name Ag7558

Primers		Length	Start Position	SEQ ID No
Forward	5'-tcgaagggcacatcatca-3'	18	4319	417
Probe	TET-5'-tgcctctgtccctggctgaaattctc-3'-TAMRA	26	4348	418
Reverse	5'-tgaagatgctggtcttcctca-3'	21	4400	419

Table AQB. CNS_neurodegeneration_v1.0

Tissue Name	Rel. Exp.(%) Ag7558, Run 308750599	issue Name	Rel. Exp.(%) Ag7558, Run 308750599
AD 1 Hippo	4.2	Control (Path) 3 Temporal Ctx	3.3
AD 2 Hippo	16.4	Control (Path) 4 Temporal Ctx	50.3
AD 3 Hippo	1.7	AD 1 Occipital Ctx	11.1
AD 4 Hippo	11.3	AD 2 Occipital Ctx (Missing)	0.0
AD 5 Hippo	76.3	AD 3 Occipital Ctx	2.3
AD 6 Hippo	38.7	AD 4 Occipital Ctx	19.8
Control 2 Hippo	17.8	AD 5 Occipital Ctx	45.4
Control 4 Hippo	3.9	AD 6 Occipital Ctx	21.2
Control (Path) 3 Hippo	1.0	Control 1 Occipital Ctx	0.9
AD 1 Temporal Ctx	7.6	Control 2 Occipital Ctx	82.4

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AD 2 Temporal Ctx	24.5	Control 3 Occipital Ctx	13.4
AD 3 Temporal Ctx	4.0	Control 4 Occipital Ctx	0.0
AD 4 Temporal Ctx	32.3	Control (Path) 1 Occipital Ctx	**100.0**
AD 5 Inf Temporal Ctx	78.5	Control (Path) 2 Occipital Ctx	17.1
AD 5 Sup Temporal Ctx	25.3	Control (Path) 3 Occipital Ctx	0.0
AD 6 Inf Temporal Ctx	39.2	Control (Path) 4 Occipital Ctx	31.9
AD 6 Sup Temporal Ctx	71.7	Control 1 Parietal Ctx	1.8
Control 1 Temporal Ctx	4.3	Control 2 Parietal Ctx	36.9
Control 2 Temporal Ctx	33.2	Control 3 Parietal Ctx	21.5
Control 3 Temporal Ctx	13.8	Control (Path) 1 Parietal Ctx	87.1
Control 3 Temporal Ctx	2.5	Control (Path) 2 Parietal Ctx	41.5
Control (Path) 1 Temporal Ctx	55.9	Control (Path) 3 Parietal Ctx	3.7
Control (Path) 2 Temporal Ctx	65.1	Control (Path) 4 Parietal Ctx	79.0

Table AQC. Panel 5 Islet

5

Tissue Name	Rel. Exp.(%) Ag7558, Run 312000203	Tissue Name	Rel. Exp.(%) Ag7558, Run 312000203
97457_Patient-02go_adipose	0.0	94709_Donor 2 AM - A_adipose	0.0
97476_Patient-07sk_skeletal muscle	0.0	94710_Donor 2 AM - B_adipose	0.0
97477_Patient-07ut_uterus	0.0	94711_Donor 2 AM - C_adipose	0.0
97478_Patient-07pl_placenta	0.0	94712_Donor 2 AD - A_adipose	0.0
99167_Bayer Patient 1	**100.0**	94713_Donor 2 AD - B_adipose	0.0
97482_Patient-08ut_uterus	0.0	94714_Donor 2 AD - C_adipose	0.0
97483_Patient-08pl_placenta	0.0	94742_Donor 3 U - A_Mesenchymal Stem Cells	0.0
97486_Patient-09sk_skeletal muscle	0.0	94743_Donor 3 U - B_Mesenchymal Stem Cells	0.0
97487_Patient-09ut_uterus	0.0	94730_Donor 3 AM - A_adipose	0.0
97488_Patient-09pl_placenta	0.0	94731_Donor 3 AM - B_adipose	0.0
97492_Patient-10ut_uterus	0.0	94732_Donor 3 AM - C_adipose	0.0
97493_Patient-10pl_placenta	0.0	94733_Donor 3 AD - A_adipose	0.0
97495_Patient-11go_adipose	0.0	94734_Donor 3 AD - B_adipose	0.0
97496_Patient-11sk_skeletal muscle	0.0	94735_Donor 3 AD - C_adipose	0.0
97497_Patient-11ut_uterus	0.0	77138_Liver_HepG2untreated	0.0
97498_Patient-11pl_placenta	0.0	73556_Heart_Cardiac stromal cells (primary)	0.0
97500_Patient-12go_adipose	0.0	81735_Small Intestine	0.0

97501_Patient-12sk_skeletal muscle	0.0	72409_Kidney_Proximal Convoluted Tubule	0.0
97502_Patient-12ut_uterus	0.0	82685_Small intestine_Duodenum	0.0
97503_Patient-12pl_placenta	0.0	90650_Adrenal_Adrenocortical adenoma	0.0
94721_Donor 2 U - A_Mesenchymal Stem Cells	0.0	72410_Kidney_HRCE	0.0
94722_Donor 2 U - B_Mesenchymal Stem Cells	0.0	72411_Kidney_HRE	0.0
94723_Donor 2 U - C_Mesenchymal Stem Cells	0.0	73139_Uterus_Uterine smooth muscle cells	0.0

CNS_neurodegeneration_v1.0 Summary: Ag7558 Highest expression of this gene is seen in the occipital cortex of a control patient (CT=33). This panel does not show differential expression of this gene in Alzheimer's disease. However, this profile does show the expression of this gene at low levels in the brain. Therefore, therapeutic modulation of the expression or function of this gene may be useful in the treatment of neurological disorders, such as Alzheimer's disease, Parkinson's disease, schizophrenia, multiple sclerosis, stroke and epilepsy.

Panel 4.1D Summary: Ag7558 Expression of this gene is low/undetectable in all samples on this panel (CTs>35).

Panel 5 Islet Summary: Ag7558 Expression of this gene is limited to pancreatic islet cells (CT=34.6). This gene codes for a variant of SUR1. SUR1 is a subunit of the pancreatic beta cell K+ channel that regulates insulin release in glucose-stimulated cells. Thus, therapeutic modulation of SUR1 variant encoded by this gene may be used as a treatment for the enhancement of insulin secretion in Type 2 diabetes.

AR. CG93659-03: MITOGEN-ACTIVATED PROTEIN KINASE KINASE KINASE 9.

Expression of gene CG93659-03 was assessed using the primer-probe set Ag4828, described in Table ARA. Results of the RTQ-PCR runs are shown in Tables ARB and ARC.

Table ARA. Probe Name Ag4828

Primers		Length	Start Position	SEQ ID No

Forward	5'-gaggaatctgagatgctcaaga-3'	22	1275	420
Probe	TET-5'-caacgctctctctacatcgacctcgg-3'-TAMRA	26	1299	421
Reverse	5'-tccccgaacaagattgaagt-3'	20	1339	422

Table ARB. General screening panel v1.4

Tissue Name	Rel. Exp.(%) Ag4828, Run 217081802	issue Name	Rel. Exp.(%) Ag4828, Run 217081802
Adipose	53.6	Renal ca. TK-10	10.6
Melanoma* Hs688(A).T	15.5	Bladder	31.9
Melanoma* Hs688(B).T	17.4	Gastric ca. (liver met.) NCI-N87	36.3
Melanoma* M14	3.5	Gastric ca. KATO III	12.2
Melanoma* LOXIMVI	3.2	Colon ca. SW-948	5.4
Melanoma* SK-MEL-5	0.9	Colon ca. SW480	25.0
Squamous cell carcinoma SCC-4	7.0	Colon ca.* (SW480 met) SW620	2.5
Testis Pool	4.7	Colon ca. HT29	14.3
Prostate ca.* (bone met) PC-3	6.3	Colon ca. HCT-116	2.1
Prostate Pool	3.9	Colon ca. CaCo-2	15.9
Placenta	39.0	Colon cancer tissue	39.8
Uterus Pool	9.0	Colon ca. SW1116	3.4
Ovarian ca. OVCAR-3	15.7	Colon ca. Colo-205	8.8
Ovarian ca. SK-OV-3	46.3	Colon ca. SW-48	5.4
Ovarian ca. OVCAR-4	7.1	Colon Pool	16.2
Ovarian ca. OVCAR-5	30.6	Small Intestine Pool	9.3
Ovarian ca. IGROV-1	14.1	Stomach Pool	17.3
Ovarian ca. OVCAR-8	2.7	Bone Marrow Pool	7.0
Ovary	4.5	Fetal Heart	2.9
Breast ca. MCF-7	**100.0**	Heart Pool	7.9
Breast ca. MDA-MB-231	9.2	Lymph Node Pool	15.2
Breast ca. BT 549	73.2	Fetal Skeletal Muscle	1.7
Breast ca. T47D	66.0	Skeletal Muscle Pool	9.8
Breast ca. MDA-N	0.9	Spleen Pool	45.7
Breast Pool	24.1	Thymus Pool	15.9
Trachea	18.0	CNS cancer (glio/astro) U87-MG	7.6
Lung	6.7	CNS cancer (glio/astro) U-118-MG	7.9
Fetal Lung	68.3	CNS cancer (neuro;met) SK-N-AS	2.6
Lung ca. NCI-N417	0.2	CNS cancer (astro) SF-539	2.3
Lung ca. LX-1	11.8	CNS cancer (astro) SNB-75	14.1
Lung ca. NCI-H146	0.0	CNS cancer (glio) SNB-19	11.1

Lung ca. SHP-77	0.1	CNS cancer (glio) SF-295	3.9
Lung ca. A549	36.6	Brain (Amygdala) Pool	2.7
Lung ca. NCI-H526	0.0	Brain (cerebellum)	1.4
Lung ca. NCI-H23	13.4	Brain (fetal)	4.9
Lung ca. NCI-H460	17.6	Brain (Hippocampus) Pool	3.7
Lung ca. HOP-62	13.2	Cerebral Cortex Pool	3.5
Lung ca. NCI-H522	2.1	Brain (Substantia nigra) Pool	2.7
Liver	1.0	Brain (Thalamus) Pool	4.5
Fetal Liver	2.8	Brain (whole)	4.5
Liver ca. HepG2	8.1	Spinal Cord Pool	3.8
Kidney Pool	31.4	Adrenal Gland	9.5
Fetal Kidney	7.7	Pituitary gland Pool	1.4
Renal ca. 786-0	10.9	Salivary Gland	2.5
Renal ca. A498	5.2	Thyroid (female)	7.7
Renal ca. ACHN	2.5	Pancreatic ca. CAPAN2	34.4
Renal ca. UO-31	14.9	Pancreas Pool	19.6

Table ARC. Panel 5D

5

Tissue Name	Rel. Exp. %) Ag4828, Run 219436967	Tissue Name	Rel. Exp.(%) Ag4828, Run 219436967
97457_Patient-02go_adipose	33.9	94709_Donor 2 AM - A_adipose	10.8
97476_Patient-07sk_skeletal muscle	33.4	94710_Donor 2 AM - B_adipose	9.3
97477_Patient-07ut_uterus	59.5	94711_Donor 2 AM - C_adipose	3.0
97478_Patient-07pl_placenta	39.8	94712_Donor 2 AD - A_adipose	13.7
97481_Patient-08sk_skeletal muscle	25.9	94713_Donor 2 AD - B_adipose	10.0
97482_Patient-08ut_uterus	19.8	94714_Donor 2 AD - C_adipose	6.7
97483_Patient-08pl_placenta	41.5	94742_Donor 3 U - A_Mesenchymal Stem Cells	4.7
97486_Patient-09sk_skeletal muscle	6.5	94743_Donor 3 U - B_Mesenchymal Stem Cells	2.8
97487_Patient-09ut_uterus	8.1	94730_Donor 3 AM - A_adipose	6.3
97488_Patient-09pl_placenta	38.4	94731_Donor 3 AM - B_adipose	2.4
97492_Patient-10ut_uterus	30.6	94732_Donor 3 AM - C_adipose	2.2
97493_Patient-10pl_placenta	72.7	94733_Donor 3 AD - A_adipose	10.2
97495_Patient-11go_adipose	100.0	94734_Donor 3 AD - B_adipose	5.5
97496_Patient-11sk_skeletal muscle	5.8	94735_Donor 3 AD - C_adipose	4.7
97497_Patient-11ut_uterus	20.6	77138_Liver_HepG2untreated	14.4

97498_Patient-11pl_placenta	50.0	73556_Heart_Cardiac stromal cells (primary)	1.9
97500_Patient-12go_adipose	82.4	81735_Small Intestine	17.2
97501_Patient-12sk_skeletal muscle	19.2	72409_Kidney_Proximal Convoluted Tubule	0.9
97502_Patient-12ut_uterus	23.7	82685_Small intestine_Duodenum	19.1
97503_Patient-12pl_placenta	57.0	90650_Adrenal_Adrenocortical adenoma	8.8
94721_Donor 2 U - A_Mesenchymal Stem Cells	1.6	72410_Kidney_HRCE	7.6
94722_Donor 2 U - B_Mesenchymal Stem Cells	3.0	72411_Kidney_HRE	13.5
94723_Donor 2 U - C_Mesenchymal Stem Cells	2.1	73139_Uterus_Uterine smooth muscle cells	2.0

General_screening_panel_v1.4 Summary: Ag4828 Highest expression of this gene is detected in a breast cancer MCF-7 cell line(CT=27.6). Interestingly, this gene is expressed at much higher levels in fetal (CT=28) when compared to adult lung (CT=31). This observation suggests that expression of this gene can be used to distinguish fetal from adult lung. In addition, the relative overexpression of this gene in fetal lung suggests that the protein product may enhance lung growth or development in the fetus and thus may also act in a regenerative capacity in the adult. Therefore, therapeutic modulation of the protein encoded by this gene could be useful in treatment of lung related diseases.

In addition significant expression of this gene is found in a number of cancer (pancreatic, CNS, colon, lung, breast, ovary, prostate, melanoma) cell lines. Therefore, therapeutic modulation of the activity of this gene or its protein product, through the use of small molecule drugs, might be beneficial in the treatment of these cancers.

Among tissues with metabolic or endocrine function, this gene is expressed at high to moderate levels in pancreas, adipose, adrenal gland, thyroid, skeletal muscle, heart, fetal liver and the gastrointestinal tract. Therefore, therapeutic modulation of the activity of this gene may prove useful in the treatment of endocrine/metabolically related diseases, such as obesity and diabetes.

This gene encodes a protein that is homologous to mitogen-activated protein kinase kinase kinase 8 (MAP3K8)(COT proto-oncogene serine/threonine-protein kinase) (C-COT) (Cancer osaka thyroid oncogene). COT is able to enhance the TNF alpha production and to activate NF-kB. Both events are connected with insulin resistance and type II diabetes (1,

2, 3). Inhibition of COT kinase would prevent overproduction of TNF alpha and activation of NF-kB, thus improving insulin resistance and diabetes.

In addition, this gene is expressed at high levels in all regions of the central nervous system examined, including amygdala, hippocampus, substantia nigra, thalamus, cerebellum, cerebral cortex, and spinal cord. Recently, MKK6, a related protein, has been shown to associated with Alzheimer's disease (4). Therefore, based on the homology of this protein to MKK6 and the presence of this gene in the brain, we predict that this putative MAP3K8 may play a role in central nervous system disorders such as Alzheimer's disease, Parkinson's disease, epilepsy, multiple sclerosis, schizophrenia and depression.

References:

1. Ballester A, Velasco A, Tobena R, Alemany S. Cot kinase activates tumor necrosis factor-alpha gene expression in a cyclosporin A-resistant manner. J. Biol. Chem. 1998. 273, 14099-106. PMID: 9603908.

2. Bierhaus A, Schiekofer S, Schwaninger M, Andrassy M, Humpert PM, Chen J, Hong M, Luther T, Henle T, Kloting I, Morcos M, Hofmann M, Tritschler H, Weigle B, Kasper M, Smith M, Perry G, Schmidt AM, Stern DM, Haring HU, Schleicher E, Nawroth PP. Diabetes-associated sustained activation of the transcription factor nuclear factor-kappaB. Diabetes, 2001 50, 2792-808. PMID: 11723063.

3. Belich MP, Salmeron A, Johnston LH, Ley SC. TPL-2 kinase regulates the proteolysis of the NF-kappaB-inhibitory protein NF-kappaB1 p105. Nature. 1999 397, 363-8.PMID: 9950430.

4. Zhu X, Rottkamp CA, Hartzler A, Sun Z, Takeda A, Boux H, Shimohama S, Perry G, Smith MA. (2001) Activation of MKK6, an upstream activator of p38, in Alzheimer's disease. J Neurochem 79(2):311-8

Panel 5D Summary: Ag4828 Highest expression of this gene is detected in adipose tissue (CT=29). Low to moderate expression of this gene is seen in wide range of samples used in this panel including adipose, skeletal muscle, uterus, and placenta. This wide spread expression of this gene in tissues with metabolic or endocrine function, suggests that this gene plays a role in endocrine/metabolically related diseases, such as obesity and diabetes.

This gene encodes a MAP3K8-like protein. Recently, activation of MAP kinase, ERK, a related protein, by modified LDL in vascular smooth muscle cells has been

implicated in the development of atherosclerosis in diabetes (Ref. 1). Therefore, this putative MAP3K8 may also play a role in the development of this disease. Therefore, therapeutic modulation of the activity of this gene or its protein product, through the use of small molecule drugs, might be beneficial in the treatment of artherosclerosis and diabetes.

5   References:

1. Velarde V, Jenkins AJ, Christopher J, Lyons TJ, Jaffa AA. (2001) Activation of MAPK by modified low-density lipoproteins in vascular smooth muscle cells. J Appl Physiol 91(3):1412-20

AS. CG94521-02 and CG94521-03: CYTOPLASMIC
10   GLYCEROL-3-PHOSPHATE DEHYDROGENASE [NAD+].

Expression of gene CG94521-02 and CG94521-03 was assessed using the primer-probe set Ag3924, described in Table ASA. Results of the RTQ-PCR runs are shown in Tables ASB, ASC, ASD, ASE and ASF. Please note that these sequences represent full-length physical clones.

15   **Table ASA. Probe Name Ag3924**

Primers		Length	Start Position	SEQ ID No
Forward	5'-actgggaagaccattgaagagt-3'	22	197	423
Probe	TET-5'-aaaagctccaaggaccgcagacttct-3'-TAMRA	26	147	424
Reverse	5'-gtttgaggatgcggtacactt-3'	21	122	425

20   **Table ASB. CNS_neurodegeneration_v1.0**

Tissue Name	Rel. Exp.(%) Ag3924, Run 212343350	issue Name	Rel. Exp.(%) Ag3924, Run 212343350
AD 1 Hippo	8.4	Control (Path) 3 Temporal Ctx	6.0
AD 2 Hippo	21.9	Control (Path) 4 Temporal Ctx	2.8
AD 3 Hippo	8.4	AD 1 Occipital Ctx	14.4
AD 4 Hippo	7.5	AD 2 Occipital Ctx (Missing)	0.0
AD 5 hippo	92.7	AD 3 Occipital Ctx	4.8
AD 6 Hippo	24.5	AD 4 Occipital Ctx	14.0

Control 2 Hippo	25.7	AD 5 Occipital Ctx	14.0
Control 4 Hippo	7.3	AD 6 Occipital Ctx	55.5
Control (Path) 3 Hippo	8.8	Control 1 Occipital Ctx	6.1
AD 1 Temporal Ctx	8.3	Control 2 Occipital Ctx	47.3
AD 2 Temporal Ctx	23.8	Control 3 Occipital Ctx	9.8
AD 3 Temporal Ctx	4.2	Control 4 Occipital Ctx	4.5
AD 4 Temporal Ctx	15.1	Control (Path) 1 Occipital Ctx	64.6
AD 5 Inf Temporal Ctx	100.0	Control (Path) 2 Occipital Ctx	8.6
AD 5 SupTemporal Ctx	32.3	Control (Path) 3 Occipital Ctx	3.9
AD 6 Inf Temporal Ctx	39.0	Control (Path) 4 Occipital Ctx	15.8
AD 6 Sup Temporal Ctx	33.2	Control 1 Parietal Ctx	5.0
Control 1 Temporal Ctx	4.5	Control 2 Parietal Ctx	40.3
Control 2 Temporal Ctx	44.4	Control 3 Parietal Ctx	14.6
Control 3 Temporal Ctx	11.1	Control (Path) 1 Parietal Ctx	70.7
Control 4 Temporal Ctx	4.4	Control (Path) 2 Parietal Ctx	15.5
Control (Path) 1 Temporal Ctx	49.0	Control (Path) 3 Parietal Ctx	4.9
Control (Path) 2 Temporal Ctx	29.9	Control (Path) 4 Parietal Ctx	39.5

Table ASC. General_screening_panel_v1.4

5

Tissue Name	Rel. Exp.(%) Ag3924, Run 219515221	Tissue Name	Rel. Exp.(%) Ag3924, Run 219515221
Adipose	14.0	Renal ca. TK-10	7.1
Melanoma* Hs688(A).T	3.6	Bladder	8.1
Melanoma* Hs688(B).T	4.9	Gastric ca. (liver met.) NCI-N87	7.7
Melanoma* M14	15.1	Gastric ca. KATO III	17.4
Melanoma* LOXIMVI	6.2	Colon ca. SW-948	25.5
Melanoma* SK-MEL-5	37.6	Colon ca. SW480	28.3
Squamous cell carcinoma SCC-4	1.1	Colon ca.* (SW480 met) SW620	6.6
Testis Pool	6.3	Colon ca. HT29	4.1
Prostate ca.* (bone met) PC-3	47.0	Colon ca. HCT-116	25.0
Prostate Pool	18.6	Colon ca. CaCo-2	6.9
Placenta	6.3	Colon cancer tissue	7.6
Uterus Pool	5.1	Colon ca. SW1116	5.2
Ovarian ca. OVCAR-3	11.3	Colon ca. Colo-205	2.6
Ovarian ca. SK-OV-3	6.8	Colon ca. SW-48	4.4
Ovarian ca. OVCAR-4	12.2	Colon Pool	9.9
Ovarian ca. OVCAR-5	17.9	Small Intestine Pool	9.3
Ovarian ca. IGROV-1	8.2	Stomach Pool	5.2
Ovarian ca. OVCAR-8	3.5	Bone Marrow Pool	4.9

Ovary	9.6	Fetal Heart	26.1
Breast ca. MCF-7	100.0	Heart Pool	23.7
Breast ca. MDA-MB-231	11.4	Lymph Node Pool	8.7
Breast ca. BT 549	11.4	Fetal Skeletal Muscle	11.2
Breast ca. T47D	40.9	Skeletal Muscle Pool	62.0
Breast ca. MDA-N	11.7	Spleen Pool	9.7
Breast Pool	8.3	Thymus Pool	5.8
Trachea	15.4	CNS cancer (glio/astro) U87-MG	18.2
Lung	2.8	CNS cancer (glio/astro) U-118-MG	11.3
Fetal Lung	21.8	CNS cancer (neuro;met) SK-N-AS	6.6
Lung ca. NCI-N417	13.4	CNS cancer (astro) SF-539	4.0
Lung ca. LX-1	8.2	CNS cancer (astro) SNB-75	21.9
Lung ca. NCI-H146	4.5	CNS cancer (glio) SNB-19	7.6
Lung ca. SHP-77	13.3	CNS cancer (glio) SF-295	24.0
Lung ca. A549	16.6	Brain (Amygdala) Pool	11.4
Lung ca. NCI-H526	2.4	Brain (cerebellum)	10.2
Lung ca. NCI-H23	2.0	Brain (fetal)	27.2
Lung ca. NCI-H460	2.9	Brain (Hippocampus) Pool	11.6
Lung ca. HOP-62	6.6	Cerebral Cortex Pool	17.2
Lung ca. NCI-H522	14.3	Brain (Substantia nigra) Pool	10.4
Liver	0.3	Brain (Thalamus) Pool	18.9
Fetal Liver	1.1	Brain (whole)	17.7
Liver ca. HepG2	3.4	Spinal Cord Pool	14.3
Kidney Pool	26.4	Adrenal Gland	37.9
Fetal Kidney	6.7	Pituitary gland Pool	5.0
Renal ca. 786-0	3.0	Salivary Gland	11.1
Renal ca. A498	1.4	Thyroid (female)	17.0
Renal ca. ACHN	2.5	Pancreatic ca. CAPAN2	2.8
Renal ca. UO-31	10.1	Pancreas Pool	13.3

Table ASD. Panel 4.1D

5

Tissue Name	Rel. Exp.(% Ag3924, Run 170552351	Tissue Name	Rel. Exp.(%) Ag3924, Run 170552351
Secondary Th1 act	33.9	HUVEC IL-1beta	19.6
Secondary Th2 act	35.4	HUVEC IFN gamma	32.3
Secondary Tr1 act	29.3	HUVEC TNF alpha + IFN gamma	8.6
Secondary Th1 rest	14.8	HUVEC TNF alpha + IL4	19.1
Secondary Th2 rest	23.7	HUVEC IL-11	17.2
Secondary Tr1 rest	15.8	Lung Microvascular EC none	16.8

Primary Th1 act	31.0	Lung Microvascular EC TNFalpha + IL-1beta	11.0
Primary Th2 act	33.7	Microvascular Dermal EC none	27.7
Primary Tr1 act	33.9	Microsvasular Dermal EC TNFalpha + IL-1beta	8.6
Primary Th1 rest	27.4	Bronchial epithelium TNFalpha + IL1beta	6.7
Primary Th2 rest	15.3	Small airway epithelium none	4.7
Primary Tr1 rest	34.2	Small airway epithelium TNFalpha + IL-1beta	4.0
CD45RA CD4 lymphocyte act	17.4	Coronery artery SMC rest	8.1
CD45RO CD4 lymphocyte act	28.3	Coronery artery SMC TNFalpha + IL-1beta	4.4
CD8 lymphocyte act	24.1	Astrocytes rest	16.4
Secondary CD8 lymphocyte rest	18.2	Astrocytes TNFalpha + IL-1beta	11.9
Secondary CD8 lymphocyte act	15.2	KU-812 (Basophil) rest	37.1
CD4 lymphocyte none	12.8	KU-812 (Basophil) PMA/ionomycin	35.6
2ry Th1/Th2/Tr1_anti-CD95 CH11	21.0	CCD1106 (Keratinocytes) none	9.5
LAK cells rest	17.8	CCD1106 (Keratinocytes) TNFalpha + IL-1beta	4.8
LAK cells IL-2	26.6	Liver cirrhosis	14.4
LAK cells IL-2+IL-12	17.8	NCI-H292 none	42.9
LAK cells IL-2+IFN gamma	17.8	NCI-H292 IL-4	57.0
LAK cells IL-2+ IL-18	32.5	NCI-H292 IL-9	81.2
LAK cells PMA/ionomycin	7.9	NCI-H292 IL-13	60.7
NK Cells IL-2 rest	35.6	NCI-H292 IFN gamma	39.0
Two Way MLR 3 day	17.3	HPAEC none	21.2
Two Way MLR 5 day	17.1	HPAEC TNF alpha + IL-1 beta	13.4
Two Way MLR 7 day	**100.0**	Lung fibroblast none	18.0
PBMC rest	15.6	Lung fibroblast TNF alpha + IL-1 beta	6.0
PBMC PWM	16.5	Lung fibroblast IL-4	19.5
PBMC PHA-L	13.8	Lung fibroblast IL-9	30.8
Ramos (B cell) none	64.6	Lung fibroblast IL-13	22.2
Ramos (B cell) ionomycin	70.2	Lung fibroblast IFN gamma	20.0
B lymphocytes PWM	23.8	Dermal fibroblast CCD1070 rest	12.5
B lymphocytes CD40L and IL-4	17.0	Dermal fibroblast CCD1070 TNF alpha	30.1
EOL-1 dbcAMP	10.8	Dermal fibroblast CCD1070 IL-1 beta	5.4
EOL-1 dbcAMP PMA/ionomycin	2.2	Dermal fibroblast IFN gamma	8.2
Dendritic cells none	13.6	Dermal fibroblast IL-4	17.8
Dendritic cells LPS	4.5	Dermal Fibroblasts rest	20.0

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Dendritic cells anti-CD40	21.6	Neutrophils TNFa+LPS	21.37
Monocytes rest	19.8	Neutrophils rest	3.6
Monocytes LPS	3.0	Colon	35.6
Macrophages rest	14.9	Lung	27.7
Macrophages LPS	1.7	Thymus	27.7
HUVEC none	16.7	Kidney	66.4
HUVEC starved	17.7		

Table ASE. Panel 5 Islet

5

Tissue Name	Rel. Exp.(% Ag3924, Run 268363571	Tissue Name	Rel. Exp.(%) Ag3924, Run 268363571
97457_Patient-02go_adipose	18.2	94709_Donor 2 AM - A_adipose	19.6
97476_Patient-07sk_skeletal muscle	10.6	94710_Donor 2 AM - B_adipose	13.3
97477_Patient-07ut_uterus	10.2	94711_Donor 2 AM - C_adipose	11.0
97478_Patient-07pl_placenta	17.0	94712_Donor 2 AD - A_adipose	9.5
99167_Bayer Patient 1	6.5	94713_Donor 2 AD - B_adipose	21.9
97482_Patient-08ut_uterus	6.8	94714_Donor 2 AD - C_adipose	16.7
97483_Patient-08pl_placenta	11.7	94742_Donor 3 U - A_Mesenchymal Stem Cells	1.8
97486_Patient-09sk_skeletal muscle	10.6	94743_Donor 3 U - B_Mesenchymal Stem Cells	1.7
97487_Patient-09ut_uterus	12.0	94730_Donor 3 AM - A_adipose	19.6
97488_Patient-09pl_placenta	15.4	94731_Donor 3 AM - B_adipose	12.5
97492_Patient-10ut_uterus	12.9	94732_Donor 3 AM - C_adipose	12.2
97493_Patient-10pl_placenta	29.5	94733_Donor 3 AD - A_adipose	10.2
97495_Patient-11go_adipose	17.9	94734_Donor 3 AD - B_adipose	9.2
97496_Patient-11sk_skeletal muscle	70.7	94735_Donor 3 AD - C_adipose	8.9
97497_Patient-11ut_uterus	18.8	77138_Liver_HepG2untreated	11.1
97498_Patient-11pl_placenta	10.3	73556_Heart_Cardiac stromal cells (primary)	5.2
97500_Patient-12go_adipose	31.9	81735_Small Intestine	15.9
97501_Patient-12sk_skeletal muscle	100.0	72409_Kidney_Proximal Convoluted Tubule	6.5
97502_Patient-12ut_uterus	23.8	82685_Small intestine_Duodenum	17.0
97503_Patient-12pl_placenta	8.7	90650_Adrenal_Adrenocortical adenoma	14.4
94721_Donor 2 U - A_Mesenchymal Stem Cells	3.9	72410_Kidney_HRCE	11.5

| 94722_Donor 2 U - B_Mesenchymal Stem Cells | 2.8 | 72411_Kidney_HRE | 3.4 |
| 94723_Donor 2 U - C_Mesenchymal Stem Cells | 4.8 | 73139_Uterus_Uterine smooth muscle cells | 2.1 |

Table ASF. general oncology screening panel v 2.4

Tissue Name	Rel. Exp.(%) Ag3924, Run 268143856	Tissue Nme	Rel. Exp.(%) Ag3924, Run 268143856
Colon cancer 1	60.3	Bladder NAT 2	3.3
Colon NAT 1	29.7	Bladder NAT 3	2.4
Colon cancer 2	26.1	Bladder NAT 4	25.7
Colon NAT 2	60.7	Prostate adenocarcinoma 1	**100.0**
Colon cancer 3	88.9	Prostate adenocarcinoma 2	14.6
Colon NAT 3	88.9	Prostate adenocarcinoma 3	86.5
Colon malignant cancer 4	98.6	Prostate adenocarcinoma 4	34.9
Colon NAT 4	29.5	Prostate NAT 5	26.2
Lung cancer 1	17.3	Prostate adenocarcinoma 6	24.5
Lung NAT 1	7.9	Prostate adenocarcinoma 7	39.5
Lung cancer 2	31.9	Prostate adenocarcinoma 8	15.2
Lung NAT 2	14.8	Prostate adenocarcinoma 9	53.6
Squamous cell carcinoma 3	34.2	Prostate NAT 10	12.6
Lung NAT 3	5.0	Kidney cancer 1	12.0
Metastatic melanoma 1	28.3	Kidney NAT 1	25.9
Melanoma 2	4.8	Kidney cancer 2	53.6
Melanoma 3	12.9	Kidney NAT 2	64.6
Metastatic melanoma 4	42.6	Kidney cancer 3	12.5
Metastatic melanoma 5	70.7	Kidney NAT 3	26.6
Bladder cancer 1	9.3	Kidney cancer 4	15.0
Bladder NAT 1	0.0	Kidney NAT 4	14.6
Bladder cancer 2	17.7		

CNS_neurodegeneration_v1.0 Summary: Ag3924 This panel does not show differential expression of this gene in Alzheimer's disease. However, this profile confirms the expression of this gene at moderate levels in the brain. Please see Panel 1.4 for discussion of this gene in the central nervous system.

General_screening_panel_v1.4 Summary: Ag3924 Highest expression of this gene is seen in a breast cancer cell line (CT=25.3). This gene is ubiquitously expressed in

this panel, with high to moderate expression seen in brain, colon, gastric, lung, breast, ovarian, and melanoma cancer cell lines. This expression profile suggests a role for this gene product in cell survival and proliferation. Modulation of this gene product may be useful in the treatment of cancer.

5   Among tissues with metabolic function, this gene is expressed at moderate to high levels in pituitary, adipose, adrenal gland, pancreas, thyroid, and adult and fetal skeletal muscle, heart, and liver. This widespread expression among these tissues suggests that this gene product may play a role in normal neuroendocrine and metabolic function and that disregulated expression of this gene may contribute to neuroendocrine disorders or

10   metabolic diseases, such as obesity and diabetes. This gene encodes a novel glycerol 3-phosphate dehydrogenase (G3PD).

Similar to known cytosolic glycerol 3-phosphate dehydrogenase, this putative G3PD may contribute to glycerol synthesis and link glycolysis with TG production. This gene is highly expressed in skeletal muscle and diabetic skeletal muscle on Panel 5I.

15   Diabetic skeletal muscle has increased glycolytic activity and increased lipid content that interfere with insulin sensitivity. Inhibition of G3PD may balance disproportionate glycolysis and impair accumulation of TG in skeletal muscle. Thus, an antagonist of this novel G3PD may be beneficial for the treatment of diabetes.

This gene is also expressed at high to moderate levels in the CNS, including the

20   hippocampus, thalamus, substantia nigra, amygdala, cerebellum and cerebral cortex. Therefore, therapeutic modulation of the expression or function of this gene may be useful in the treatment of neurologic disorders, such as Alzheimer's disease, Parkinson's disease, schizophrenia, multiple sclerosis, stroke and epilepsy.

In addition, this gene is expressed at much higher levels in fetal lung tissue

25   (CT=27.5) when compared to expression in the adult counterpart (CT=30.5). Thus, expression of this gene may be used to differentiate between the fetal and adult source of this tissue.

Panel 4.1D Summary: Ag3924 Highest expression is seen in a sample derived from an MLR, where the sample was take 7 days after the reaction (CT=27.6). This gene is

30   also expressed at high to moderate levels in a wide range of cell types of significance in the immune response in health and disease. These cells include members of the T-cell, B-cell, endothelial cell, macrophage/monocyte, and peripheral blood mononuclear cell family, as well as epithelial and fibroblast cell types from lung and skin, and normal tissues

represented by colon, lung, thymus and kidney. This ubiquitous pattern of expression suggests that this gene product may be involved in homeostatic processes for these and other cell types and tissues. This pattern is in agreement with the expression profile in General_screening_panel_v1.4 and also suggests a role for the gene product in cell survival and proliferation. Therefore, modulation of the gene product with a functional therapeutic may lead to the alteration of functions associated with these cell types and lead to improvement of the symptoms of patients suffering from autoimmune and inflammatory diseases such as asthma, allergies, inflammatory bowel disease, lupus erythematosus, psoriasis, rheumatoid arthritis, and osteoarthritis.

Panel 5 Islet Summary: Ag3924 Highest expression is seen in skeletal muscle from a diabetic patient (patient 12) (CT=28). This panel confirms expression of this gene in metabolic tissues including adipose, skeletal muscle and placenta. Please see Panel 1.4 for discussion of this gene in metabolic disease.

general oncology screening panel_v_2.4 Summary: Ag3924 Highest expression is seen in a prostate cancer sample (CT=28.2). Prominent expression is also seen in melanoma samples, as well as in normal and malignant kidney, colon and lung. Thus, modulation of this gene may be useful in the treatment of prostate cancer and melanoma.

AT. CG96613-02 and CG96613-03: Splice variant of PDK1.

Expression of gene CG96613-02 and CG96613-03 was assessed using the primer-probe sets Ag1778 and Ag5110, described in Tables ATA and ATB. Results of the RTQ-PCR runs are shown in Tables ATC, ATD, ATE, ATF, ATG and ATH. Please note that probe-primer set Ag1778 is specific for CG96613-03.

Table ATA. Probe Name Ag1778

Primers		Length	Start Position	SEQ ID No
Forward	5'-gattgcccatatcacgtcttta-3'	22	1241	426
Probe	TET-5'-cgcacaatacttccaaggagacctga-3'-TAMRA	26	1263	427
Reverse	5'-gataactgcatctgtcccgtaa-3'	22	1308	428

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Table ATB. Probe Name Ag5110

Primers		Length	Start Position	SEQ ID No
Forward	5'-tgtatggcctgcaagatgat-3'	20	559	429
Probe	TET-5'-tcattcccacaatggcccagg-3'-TAMRA	21	623	430
Reverse	5'-agctctccttgtattcaatcaca-3'	23	645	431

5

Table ATC. CNS_neurodegeneration_v1.0

Tissue Name	Rel. Exp.(%) Ag1778, Run 276596797	Rel. Exp.(%) Ag5110, Run 226442922	Rel. Exp.(%) Ag5110, Run 276596798	Tissue Name	Rel. Exp.(%) Ag1778, Run 276596797	Rel. Exp.(%) Ag5110, Run 226442922	Rel. Exp.(%) Ag5110, Run 276596798
AD 1 Hippo	11.7	6.2	5.3	Control (Path) 3 Temporal Ctx	6.6	12.2	17.7
AD 2 Hippo	31.4	7.4	20.3	Control (Path) 4 Temporal Ctx	33.4	15.8	13.3
AD 3 Hippo	12.5	5.3	4.9	AD 1 Occipital Ctx	23.0	7.7	8.0
AD 4 Hippo	5.4	9.4	0.0	AD 2 Occipital Ctx (Missing)	0.0	0.0	0.0
AD 5 Hippo	82.4	79.0	45.4	AD 3 Occipital Ctx	12.2	6.2	5.8
AD 6 Hippo	54.3	88.3	70.2	AD 4 Occipital Ctx	16.3	18.0	7.0
Control 2 Hippo	17.9	18.8	19.5	AD 5 Occipital Ctx	77.9	29.9	26.2
Control 4 Hippo	13.0	19.3	13.3	AD 6 Occipital Ctx	36.9	18.9	18.8
Control (Path) 3 Hippo	11.0	7.5	16.3	Control 1 Occipital Ctx	6.2	6.8	5.4
AD 1 Temporal Ctx	20.3	14.6	11.0	Control 2 Occipital Ctx	54.0	44.8	51.4
AD 2 Temporal Ctx	29.9	16.6	21.8	Control 3 Occipital Ctx	32.3	4.9	26.8

AD 3 Temporal Ctx	11.7	8.4	17.7	Control 4 Occipital Ctx	7.5	9.2	5.3
AD 4 Temporal Ctx	20.2	5.6	19.6	Control (Path) 1 Occipital Ctx	60.3	24.7	41.8
AD 5 Inf Temporal Ctx	72.2	47.0	46.3	Control (Path) 2 Occipital Ctx	12.8	9.2	6.3
AD 5 Sup Temporal Ctx	39.5	51.1	44.1	Control (Path) 3 Occipital Ctx	5.5	0.9	0.0
AD 6 Inf Temporal Ctx	75.3	84.1	84.1	Control (Path) 4 Occipital Ctx	16.6	15.5	12.3
AD 6 Sup Temporal Ctx	**100.0**	**100.0**	**100.0**	Control 1 Parietal Ctx	10.0	10.0	3.6
Control 1 Temporal Ctx	11.2	10.4	3.9	Control 2 Parietal Ctx	46.0	57.0	27.5
Control 2 Temporal Ctx	25.3	21.6	36.3	Control 3 Parietal Ctx	23.5	18.3	16.6
Control 3 Temporal Ctx	31.2	37.9	38.2	Control (Path) 1 Parietal Ctx	78.5	39.2	52.5
Control 3 Temporal Ctx	11.7	8.4	8.8	Control (Path) 2 Parietal Ctx	23.5	12.5	14.9
Control (Path) 1 Temporal Ctx	36.6	53.6	46.7	Control (Path) 3 Parietal Ctx	9.5	13.9	5.8
Control (Path) 2 Temporal Ctx	46.0	29.7	32.5	Control (Path) 4 Parietal Ctx	46.0	58.6	39.2

Table ATD. General screening panel v1.5

5

Tissue Name	Rel. Exp.(%) Ag5110, Run 228980585	Tissue Name	Rel. Exp.(%) Ag5110, Run 228980585
Adipose	5.4	Renal ca. TK-10	11.7
Melanoma* Hs688(A).T	10.7	Bladder	12.2

Melanoma* Hs688(B).T	5.8	Gastric ca. (liver met.) NCI-N87	3.8
Melanoma* M14	19.5	Gastric ca. KATO III	10.6
Melanoma* LOXIMVI	17.3	Colon ca. SW-948	2.6
Melanoma* SK-MEL-5	29.9	Colon ca. SW480	16.6
Squamous cell carcinoma SCC-4	4.2	Colon ca.* (SW480 met) SW620	10.8
Testis Pool	9.2	Colon ca. HT29	17.0
Prostate ca.* (bone met) PC-3	48.0	Colon ca. HCT-116	6.7
Prostate Pool	0.6	Colon ca. CaCo-2	9.8
Placenta	0.5	Colon cancer tissue	7.1
Uterus Pool	2.3	Colon ca. SW1116	2.5
Ovarian ca. OVCAR-3	5.5	Colon ca. Colo-205	3.5
Ovarian ca. SK-OV-3	11.8	Colon ca. SW-48	4.7
Ovarian ca. OVCAR-4	7.9	Colon Pool	0.8
Ovarian ca. OVCAR-5	17.4	Small Intestine Pool	1.2
Ovarian ca. IGROV-1	8.7	Stomach Pool	2.2
Ovarian ca. OVCAR-8	8.2	Bone Marrow Pool	1.2
Ovary	0.3	Fetal Heart	13.0
Breast ca. MCF-7	4.3	Heart Pool	4.0
Breast ca. MDA-MB-231	25.0	Lymph Node Pool	0.9
Breast ca. BT 549	21.3	Fetal Skeletal Muscle	0.6
Breast ca. T47D	2.7	Skeletal Muscle Pool	1.7
Breast ca. MDA-N	17.2	Spleen Pool	7.5
Breast Pool	0.7	Thymus Pool	11.6
Trachea	21.9	CNS cancer (glio/astro) U87-MG	48.3
Lung	1.2	CNS cancer (glio/astro) U-118-MG	71.7
Fetal Lung	4.0	CNS cancer (neuro;met) SK-N-AS	7.2
Lung ca. NCI-N417	11.3	CNS cancer (astro) SF-539	16.6
Lung ca. LX-1	20.3	CNS cancer (astro) SNB-75	24.7
Lung ca. NCI-H146	5.5	CNS cancer (glio) SNB-19	11.0
Lung ca. SHP-77	17.7	CNS cancer (glio) SF-295	27.5
Lung ca. A549	6.9	Brain (Amygdala) Pool	2.0
Lung ca. NCI-H526	11.9	Brain (cerebellum)	5.2
Lung ca. NCI-H23	4.7	Brain (fetal)	1.0
Lung ca. NCI-H460	32.3	Brain (Hippocampus) Pool	2.0
Lung ca. HOP-62	9.7	Cerebral Cortex Pool	1.9
Lung ca. NCI-H522	12.8	Brain (Substantia nigra) Pool	1.6
Liver	0.4	Brain (Thalamus) Pool	1.7
Fetal Liver	100.0	Brain (whole)	3.0
Liver ca. HepG2	15.4	Spinal Cord Pool	1.0
Kidney Pool	1.6	Adrenal Gland	14.9
Fetal Kidney	2.2	Pituitary gland Pool	0.4
Renal ca. 786-0	10.5	Salivary Gland	6.1
Renal ca. A498	0.2	Thyroid (female)	0.5
Renal ca. ACHN	8.4	Pancreatic ca. CAPAN2	2.6

| Renal ca. UO-31 | 3.7 | Pancreas Pool | | |

Table ATE. General screening panel v1.6

5

Tissue Name	Rel. Exp.(%) Ag1778, Run 277218713	Rel. Exp.(%) Ag5110, Run 277218715	Tissue Name	Rel. Exp.(%) Ag1778, Run 277218713	Rel. Exp.(%) Ag5110, Run 277218715
Adipose	8.8	8.7	Renal ca. TK-10	31.6	13.5
Melanoma* Hs688(A).T	45.1	15.5	Bladder	23.3	14.5
Melanoma* Hs688(B).T	34.6	11.7	Gastric ca. (liver met.) NCI-N87	22.1	5.0
Melanoma* M14	29.3	11.6	Gastric ca. KATO III	9.0	15.3
Melanoma* LOXIMVI	16.6	32.1	Colon ca. SW-948	9.2	4.4
Melanoma* SK-MEL-5	23.0	36.9	Colon ca. SW480	35.8	22.5
Squamous cell carcinoma SCC-4	16.6	7.2	Colon ca.* (SW480 met) SW620	24.0	11.9
Testis Pool	8.9	8.5	Colon ca. HT29	32.1	21.5
Prostate ca.* (bone met) PC-3	100.0	50.7	Colon ca. HCT-116	17.9	9.3
Prostate Pool	5.7	1.7	Colon ca. CaCo-2	21.6	13.5
Placenta	1.6	0.3	Colon cancer tissue	3.2	10.5
Uterus Pool	3.5	3.1	Colon ca. SW1116	3.8	2.7
Ovarian ca. OVCAR-3	11.6	9.5	Colon ca. Colo-205	6.7	4.5
Ovarian ca. SK-OV-3	33.0	20.3	Colon ca. SW-48	12.1	5.2
Ovarian ca. OVCAR-4	11.4	10.7	Colon Pool	6.6	1.8
Ovarian ca. OVCAR-5	28.1	24.8	Small Intestine Pool	9.0	3.0
Ovarian ca. IGROV-1	29.1	12.7	Stomach Pool	5.6	4.5
Ovarian ca. OVCAR-8	15.9	0.1	Bone Marrow Pool	5.1	2.4
Ovary	4.4	1.6	Fetal Heart	61.6	26.4
Breast ca. MCF-7	5.9	3.6	Heart Pool	6.8	8.8
Breast ca. MDA-MB-231	79.0	34.4	Lymph Node Pool	10.4	0.8
Breast ca. BT 549	35.6	15.9	Fetal Skeletal Muscle	6.6	0.6

Breast ca. T47D	3.0	3.4	Skeletal Muscle Pool	0.9	0.7
Breast ca. MDA-N	20.7	20.9	Spleen Pool	19.2	13.0
Breast Pool	7.4	1.9	Thymus Pool	20.2	12.6
Trachea	23.8	33.7	CNS cancer (glio/astro) U87-MG	47.0	51.1
Lung	4.6	1.0	CNS cancer (glio/astro) U-118-MG	43.2	100.0
Fetal Lung	17.4	8.1	CNS cancer (neuro;met) SK-N-AS	14.1	7.7
Lung ca. NCI-N417	16.2	16.0	CNS cancer (astro) SF-539	35.1	28.3
Lung ca. LX-1	38.7	8.8	CNS cancer (astro) SNB-75	50.3	30.8
Lung ca. NCI-H146	16.7	5.9	CNS cancer (glio) SNB-19	34.4	13.1
Lung ca. SHP-77	53.2	25.9	CNS cancer (glio) SF-295	93.3	46.0
Lung ca. A549	10.9	9.9	Brain (Amygdala) Pool	7.7	2.3
Lung ca. NCI-H526	10.1	10.9	Brain (cerebellum)	24.7	5.3
Lung ca. NCI-H23	12.2	9.2	Brain (fetal)	9.7	1.3
Lung ca. NCI-H460	57.4	57.8	Brain (Hippocampus) Pool	9.7	2.8
Lung ca. HOP-62	39.0	9.7	Cerebral Cortex Pool	9.6	3.3
Lung ca. NCI-H522	19.5	13.3	Brain (Substantia nigra) Pool	6.0	2.8
Liver	1.5	0.6	Brain (Thalamus) Pool	15.3	1.9
Fetal Liver	15.1	6.0	Brain (whole)	9.5	3.3
Liver ca. HepG2	41.5	18.2	Spinal Cord Pool	5.8	2.1
Kidney Pool	9.6	2.0	Adrenal Gland	27.5	23.3
Fetal Kidney	14.7	2.6	Pituitary gland Pool	2.5	1.0
Renal ca. 786-0	14.5	11.0	Salivary Gland	9.8	10.4
Renal ca. A498	2.2	0.9	Thyroid (female)	1.5	1.9
Renal ca. ACHN	9.5	10.8	Pancreatic ca. CAPAN2	9.7	5.3
Renal ca. UO-31	13.4	4.6	Pancreas Pool	18.0	7.2

Table ATF. Panel 1.3D

Tissue Name	Rel. Exp.(% Ag1778, Run 157790405	Tissue Name	Rel. Exp.(%) Ag1778, Run 157790405
Liver adenocarcinoma	6.7	Kidney (fetal)	12.1
Pancreas	1.3	Renal ca. 786-0	6.8
Pancreatic ca. CAPAN 2	2.1	Renal ca. A498	12.2
Adrenal gland	18.7	Renal ca. RXF 393	15.0
Thyroid	2.9	Renal ca. ACHN	3.2
Salivary gland	6.2	Renal ca. UO-31	8.4
Pituitary gland	5.7	Renal ca. TK-10	3.6
Brain (fetal)	2.5	Liver	3.0
Brain (whole)	4.8	Liver (fetal)	14.7
Brain (amygdala)	6.3	Liver ca. (hepatoblast) HepG2	25.5
Brain (cerebellum)	5.4	Lung	13.7
Brain (hippocampus)	22.8	Lung (fetal)	5.3
Brain (substantia nigra)	1.1	Lung ca. (small cell) LX-1	14.5
Brain (thalamus)	3.3	Lung ca. (small cell) NCI-H69	4.9
Cerebral Cortex	14.7	Lung ca. (s.cell var.) SHP-77	36.1
Spinal cord	2.3	Lung ca. (large cell)NCI-H460	12.9
glio/astro U87-MG	21.6	Lung ca. (non-sm. cell) A549	8.1
glio/astro U-118-MG	56.3	Lung ca. (non-s.cell) NCI-H23	7.3
astrocytoma SW1783	31.2	Lung ca. (non-s.cell) HOP-62	12.8
neuro*; met SK-N-AS	30.4	Lung ca. (non-s.cl) NCI-H522	4.5
astrocytoma SF-539	22.2	Lung ca. (squam.) SW 900	1.5
astrocytoma SNB-75	12.6	Lung ca. (squam.) NCI-H596	0.7
glioma SNB-19	29.9	Mammary gland	9.7
glioma U251	22.2	Breast ca.* (pl.ef) MCF-7	4.6
glioma SF-295	20.3	Breast ca.* (pl.ef) MDA-MB-231	100.0
Heart (fetal)	35.4	Breast ca.* (pl.ef) T47D	5.1
Heart	4.5	Breast ca. BT-549	45.1
Skeletal muscle (fetal)	26.1	Breast ca. MDA-N	28.9
Skeletal muscle	3.1	Ovary	4.0
Bone marrow	13.1	Ovarian ca. OVCAR-3	4.5
Thymus	6.2	Ovarian ca. OVCAR-4	3.5
Spleen	15.5	Ovarian ca. OVCAR-5	13.4
Lymph node	16.3	Ovarian ca. OVCAR-8	3.1
Colorectal	7.9	Ovarian ca. IGROV-1	4.2
Stomach	14.5	Ovarian ca.* (ascites) SK-OV-3	13.2
Small intestine	15.5	Uterus	3.1
Colon ca. SW480	9.7	Placenta	4.3

Colon ca.* SW620(SW480 met)	10.7	Prostate	2.2
Colon ca. HT29	25.5	Prostate ca.* (bone met)PC-3	16.7
Colon ca. HCT-116	5.1	Testis	20.2
Colon ca. CaCo-2	8.1	Melanoma Hs688(A).T	7.1
Colon ca. tissue(ODO3866)	8.4	Melanoma* (met) Hs688(B).T	3.8
Colon ca. HCC-2998	12.2	Melanoma UACC-62	2.0
Gastric ca.* (liver met) NCI-N87	11.1	Melanoma M14	11.4
Bladder	8.0	Melanoma LOX IMVI	10.8
Trachea	17.7	Melanoma* (met) SK-MEL-5	5.2
Kidney	0.7	Adipose	4.9

Table ATG. Panel 4.1D

Tissue Name	Rel. Exp.(%) Ag1778, Run 276596860	Rel. Exp.(%) Ag1778, Run 276686878	Rel. Exp.(%) Ag5110, Run 226444095	Rel. Exp.(%) Ag5110, Run 276596862	Rel. Exp.(%) Ag5110, Run 276686880
Secondary Th1 act	23.5	26.8	13.9	14.9	9.0
Secondary Th2 act	28.7	28.1	11.4	14.8	17.9
Secondary Tr1 act	5.4	8.4	7.9	1.9	4.5
Secondary Th1 rest	2.9	3.8	6.3	1.0	1.5
Secondary Th2 rest	7.4	4.3	11.3	4.3	2.7
Secondary Tr1 rest	4.3	4.9	6.6	4.8	1.4
Primary Th1 act	4.5	5.6	13.9	5.0	1.8
Primary Th2 act	23.2	16.8	14.4	14.4	16.5
Primary Tr1 act	22.2	23.3	13.9	11.1	12.3
Primary Th1 rest	3.1	3.3	2.2	0.0	0.0
Primary Th2 rest	6.8	4.2	5.6	0.0	0.0
Primary Tr1 rest	2.6	3.6	10.3	0.7	0.0
CD45RA CD4 lymphocyte act	25.5	26.4	9.5	18.3	16.2
CD45RO CD4 lymphocyte act	40.1	27.2	22.1	27.9	22.4
CD8 lymphocyte act	5.1	7.4	13.1	8.1	2.4
Secondary CD8 lymphocyte rest	3.3	5.1	20.9	32.3	5.1
Secondary CD8 lymphocyte act	4.3	3.7	3.3	1.3	0.0
CD4 lymphocyte none	13.3	8.6	13.7	4.3	4.9
2ry Th1/Th2/Tr1_anti-CD95 CH11	3.2	5.2	8.1	3.1	2.4
LAK cells rest	13.2	6.7	10.1	5.6	4.6

LAK cells IL-2	9.1	8.0	11.1	6.2	3.5
LAK cells IL-2+IL-12	0.8	1.3	11.0	1.7	0.0
LAK cells IL-2+IFN gamma	9.2	8.5	12.2	4.8	7.6
LAK cells IL-2+ IL-18	6.4	5.1	15.6	3.7	12.2
LAK cells PMA/ionomycin	**100.0**	**100.0**	**100.0**	**100.0**	**100.0**
NK Cells IL-2 rest	27.5	17.8	8.7	7.1	14.7
Two Way MLR 3 day	16.8	21.2	16.3	5.1	10.7
Two Way MLR 5 day	2.9	2.7	4.2	1.7	0.0
Two Way MLR 7 day	6.2	2.6	3.4	1.9	2.6
PBMC rest	3.6	3.7	5.9	2.3	3.2
PBMC PWM	9.5	6.9	4.5	1.7	1.6
PBMC PHA-L	6.9	8.0	8.7	5.0	3.4
Ramos (B cell) none	7.7	4.2	4.7	0.6	1.4
Ramos (B cell) ionomycin	36.6	32.1	11.9	9.2	6.0
B lymphocytes PWM	11.7	4.9	6.7	4.4	4.3
B lymphocytes CD40L and IL-4	34.2	21.0	13.2	15.2	19.8
EOL-1 dbcAMP	52.1	34.4	11.0	10.8	15.6
EOL-1 dbcAMP PMA/ionomycin	9.8	6.0	3.5	1.4	5.8
Dendritic cells none	9.5	7.7	7.3	6.3	5.4
Dendritic cells LPS	5.6	5.0	6.6	1.1	2.0
Dendritic cells anti-CD40	3.6	4.2	7.0	1.3	1.5
Monocytes rest	4.9	3.1	6.9	1.2	0.0
Monocytes LPS	11.3	8.4	6.8	2.9	0.0
Macrophages rest	5.7	10.2	5.7	1.9	0.0
Macrophages LPS	3.2	3.0	5.2	0.7	3.6
HUVEC none	6.0	4.2	1.8	1.3	5.2
HUVEC starved	11.0	9.5	4.4	5.9	2.3
HUVEC IL-1beta	11.9	10.1	4.9	8.1	9.0
HUVEC IFN gamma	9.2	9.4	5.5	2.7	6.5
HUVEC TNF alpha + IFN gamma	3.8	3.6	4.1	3.5	1.8
HUVEC TNF alpha + IL4	2.7	2.8	5.5	0.0	0.0
HUVEC IL-11	4.3	5.3	3.5	3.4	0.0
Lung Microvascular EC none	25.3	23.3	7.5	6.9	6.2
Lung Microvascular EC TNFalpha + IL-1beta	9.2	7.0	7.9	2.6	2.2
Microvascular Dermal EC none	1.8	2.1	3.8	0.0	0.0
Microsvasular Dermal EC TNFalpha + IL-1beta	2.0	2.6	1.9	1.3	0.0

Bronchial epithelium TNFalpha + IL1beta	8.8	14.0	10.6	3.3	3.3
Small airway epithelium none	10.7	3.0	2.4	3.4	6.0
Small airway epithelium TNFalpha + IL-1beta	31.9	31.0	21.9	30.4	15.8
Coronery artery SMC rest	25.2	19.6	9.1	13.3	13.4
Coronery artery SMC TNFalpha + IL-1beta	27.5	19.6	5.5	7.8	15.2
Astrocytes rest	8.2	15.3	2.4	1.9	2.8
Astrocytes TNFalpha + IL-1beta	5.2	2.7	3.4	0.0	5.3
KU-812 (Basophil) rest	10.7	8.1	3.5	2.0	0.0
KU-812 (Basophil) PMA/ionomycin	37.1	25.5	11.6	8.9	5.2
CCD1106 (Keratinocytes) none	20.6	20.9	13.2	4.5	6.9
CCD1106 (Keratinocytes) TNFalpha + IL-1beta	14.1	22.7	17.8	7.7	2.3
Liver cirrhosis	11.4	8.5	7.4	1.4	1.4
NCI-H292 none	12.9	7.6	7.1	5.5	7.5
NCI-H292 IL-4	11.9	12.2	4.3	4.8	5.8
NCI-H292 IL-9	16.8	12.7	7.0	3.7	11.4
NCI-H292 IL-13	12.5	10.0	6.5	4.2	7.3
NCI-H292 IFN gamma	3.9	4.1	7.6	2.6	4.2
HPAEC none	1.7	2.9	2.6	0.0	0.0
HPAEC TNF alpha + IL-1 beta	10.6	7.2	2.9	2.7	3.3
Lung fibroblast none	31.2	24.1	4.5	8.7	5.8
Lung fibroblast TNF alpha + IL-1 beta	24.3	21.6	6.6	7.5	11.2
Lung fibroblast IL-4	6.5	1.1	1.8	3.2	4.0
Lung fibroblast IL-9	19.2	28.3	8.2	6.7	7.7
Lung fibroblast IL-13	8.2	5.1	2.9	0.0	3.6
Lung fibroblast IFN gamma	15.3	14.9	5.5	3.8	12.9
Dermal fibroblast CCD1070 rest	25.0	23.3	7.8	4.6	11.0
Dermal fibroblast CCD1070 TNF alpha	74.2	45.1	14.1	23.2	36.3
Dermal fibroblast CCD1070 IL-1 beta	23.3	22.4	4.3	3.9	5.7
Dermal fibroblast IFN gamma	3.4	3.9	2.0	0.9	0.0
Dermal fibroblast IL-4	6.8	8.2	3.3	2.6	3.0
Dermal Fibroblasts rest	11.2	7.8	2.8	2.7	3.8
Neutrophils TNFa+LPS	4.5	1.6	1.6	1.8	0.0

Neutrophils rest	28.9	31.2	12.1	15.9	20.2
Colon	2.3	1.5	2.3	0.0	2.3
Lung	2.0	2.4	3.7	0.9	1.6
Thymus	13.0	14.6	6.6	0.0	5.1
Kidney	7.9	7.5	1.7	1.1	2.8

Table ATH. general oncology screening panel v 2.4

Tissue Name	Rel. Exp.(%) Ag5110, Run 259939210	Tissue Nme	Rel. Exp.(%) Ag5110, Run 259939210
Colon cancer 1	6.5	Bladder NAT 2	0.0
Colon NAT 1	5.9	Bladder NAT 3	0.0
Colon cancer 2	6.0	Bladder NAT 4	0.0
Colon NAT 2	14.2	Prostate adenocarcinoma 1	1.2
Colon cancer 3	23.7	Prostate adenocarcinoma 2	0.0
Colon NAT 3	15.7	Prostate adenocarcinoma 3	1.6
Colon malignant cancer 4	41.5	Prostate adenocarcinoma 4	14.2
Colon NAT 4	4.2	Prostate NAT 5	0.9
Lung cancer 1	7.5	Prostate adenocarcinoma 6	0.0
Lung NAT 1	0.0	Prostate adenocarcinoma 7	0.7
Lung cancer 2	28.5	Prostate adenocarcinoma 8	0.0
Lung NAT 2	1.2	Prostate adenocarcinoma 9	3.0
Squamous cell carcinoma 3	42.3	Prostate NAT 10	0.0
Lung NAT 3	0.0	Kidney cancer 1	34.2
Metastatic melanoma 1	1.4	Kidney NAT 1	4.5
Melanoma 2	10.4	Kidney cancer 2	100.0
Melanoma 3	2.1	Kidney NAT 2	3.2
Metastatic melanoma 4	2.2	Kidney cancer 3	19.6
Metastatic melanoma 5	4.5	Kidney NAT 3	1.1
Bladder cancer 1	0.0	Kidney cancer 4	37.1
Bladder NAT 1	0.0	Kidney NAT 4	1.0
Bladder cancer 2	2.3		

CNS_neurodegeneration_v1.0 Summary: Ag1778/Ag5110 This panel confirms the expression of this gene at low levels in the brains of an independent group of individuals. However, no differential expression of this gene was detected between Alzheimer's diseased postmortem brains and those of non-demented controls in this

experiment. Please see Panel 1.5 for a discussion of this gene in treatment of central nervous system disorders.

General_screening_panel_v1.5 Summary: Ag5110 Highest expression of this gene is detected in fetal liver (CT=29.4). Interestingly, this gene is expressed at much higher levels in fetal when compared to adult liver (CT=37). This observation suggests that expression of this gene can be used to distinguish fetal from adult liver. In addition, the relative overexpression of this gene in fetal tissue suggests that the protein product may enhance liver growth or development in the fetus and thus may also act in a regenerative capacity in the adult. Therefore, therapeutic modulation of the protein encoded by this gene could be useful in treatment of liver related diseases.

Among tissues with metabolic or endocrine function, this gene is expressed at low levels in adipose, adrenal gland, heart, fetal liver and stomach. This gene codes for a splice variant of pyruvate dehydrogenase [lipoamide] kinase (PDK). Pyruvate dehydrogenase kinase (PDK) catalyzes phosphorylation and inactivation of the pyruvate dehydrogenase complex (PDC). Inactivation of PDC by increased PDK activity promotes gluconeogenesis by conserving three-carbon substrates. This helps maintain glucose levels during starvation, but is detrimental in diabetes (Huang et al., 2002, Diabetes 51(2):276-83, PMID: 11812733). Therefore, therapeutic modulation of the activity of PKD encoded by gene may be useful in the treatment of endocrine/metabolically related diseases, such as obesity and diabetes.

In addition, this gene is expressed at low levels in cerebellum and whole brain. Therefore, therapeutic modulation of this gene product may be useful in the treatment of neurological disorders such as Alzheimer's disease, Parkinson's disease, epilepsy, multiple sclerosis, schizophrenia and depression.

Moderate to low levels of expression of this gene is also seen in cluster of cancer cell lines derived from pancreatic, gastric, colon, lung, liver, renal, breast, ovarian, prostate, squamous cell carcinoma, melanoma and brain cancers. Thus, expression of this gene could be used as a marker to detect the presence of these cancers. Furthermore, therapeutic modulation of the expression or function of this gene may be effective in the treatment of pancreatic, gastric, colon, lung, liver, renal, breast, ovarian, prostate, squamous cell carcinoma, melanoma and brain cancers.

General_screening_panel_v1.6 Summary: Ag1778/Ag5110 Two experiments with different probe and primer sets are in good agreement. Highest expression of this gene

538

is detected in a prostate cancer PC3 and a brain cancer U-118-MG cell lines (CTs=31.3?.3) (CTs=25-29.8). Expression in this panel correlates with pattern seen in panel 1.5. Moderate to low levels of expression of this gene is detected in tissues with metabolic/endocrine functions such as pancreas, adipose, adrenal gland, heart, fetal liver and gastrointestinal
5   tract, in brain including cerebellum, cerebral cortex, substantia nigra and the whole brain and also in number of cancer cell lines derived from pancreatic, gastric, colon, lung, liver, renal, breast, ovarian, prostate, squamous cell carcinoma, melanoma and brain cancers. Please see panel 1.5 for further discussion on the utility of this gene.

   Panel 1.3D Summary: Ag1778 Highest expression of this gene is detected in a
10   breast cancer cell line (CT=27.4). Expression in this panel correlates with pattern seen in panel 1.5. Moderate to low levels of expression of this gene is detected in tissues with metabolic/endocrine functions such as pancreas, adrenal gland, heart, fetal liver and gastrointestinal tract, in brain including cerebellum, cerebral cortex, substantia nigra and the whole brain and also in number of cancer cell lines derived from pancreatic, gastric,
15   colon, lung, liver, renal, breast, ovarian, prostate, squamous cell carcinoma, melanoma and brain cancers. Please see panel 1.5 for further discussion of this gene.

   Panel 4.1D Summary: Ag1778/Ag5110 Five experiments with the two different probe-primer sets are in good agreement. Highest expression of this gene is detected in PMA/ionomycin treated LAK cells. These cells are involved in tumor immunology and cell
20   clearance of virally and bacterial infected cells as well as tumors. Therefore, modulation of the function of the protein encoded by this gene through the application of a small molecule drug or antibody may alter the functions of these cells and lead to improvement of symptoms associated with these conditions.

   Low levels of expression of this gene is also seen in naive and memory T cells,
25   resting secondary CD8 lymphocytes, cytokine activated small airway epithelium, and resting neutrophils. Therefore, therapeutic modulation of this gene or its protein product may be useful in the treatment of Therefore, therapeutic modulation of this gene product may ameliorate symptoms/conditions associated with autoimmune and inflammatory disorders including psoriasis, allergy, asthma, inflammatory bowel disease, rheumatoid
30   arthritis and osteoarthritis

   general oncology screening panel_v_2.4 Summary: Ag5110 Highest expression of this gene is detected in kidney cancer (CT=32). Low levels of expression of this gene is also seen in colon, lung, prostate and kidney cancer. Higher levels of expression of this

gene is associated with cancer as compared to corresponding normal tissue. Therefore, expression of this gene may be used as diagnostic marker for the detection of these cancers. Furthermore, therapeutic modulation of this gene or its protein product may be useful in the treatment of colon, lung, prostate and kidney cancers.

5   **AU. CG96736-01: Neutral amino acid transporter B.**

Expression of gene CG96736-01 was assessed using the primer-probe sets Ag3788 and Ag4075, described in Tables AUA and AUB. Results of the RTQ-PCR runs are shown in Tables AUC, AUD, AUE, AUF, AUG, AUH, AUI, AUJ and AUK.

Table AUA. Probe Name Ag3788

10

Primers		Length	Start Position	SEQ ID No
Forward	5'-cgagaaatatcttcccttccaa-3'	22	1182	432
Probe	TET-5'-tgtcagcagcctttcgctcatactct-3'-TAMRA	26	1209	433
Reverse	5'-ttccggtgatattcctctcttc-3'	22	1244	434

Table AUB. Probe Name Ag4075

15

Primers		Length	Start Position	SEQ ID No
Forward	5'-cgagaaatatcttcccttccaa-3'	22	1182	435
Probe	TET-5'-tgtcagcagcctttcgctcatactct-3'-TAMRA	26	1209	436
Reverse	5'-ttccggtgatattcctctcttc-3'	22	1244	437

Table AUC. AI comprehensive panel_v1.0

20

Tissue Name	Rel. Exp.(%) Ag4075, Run 226203371	Tissue Name	Rel. Exp.(%) Ag4075, Run 226203371
110967 COPD-F	6.0	112427 Match Control Psoriasis-F	12.3
110980 COPD-F	9.9	112418 Psoriasis-M	3.6
110968 COPD-M	6.6	112723 Match Control Psoriasis-M	6.3
110977 COPD-M	0.0	112419 Psoriasis-M	6.5

110989 Emphysema-F	8.7	112424 Match Control Psoriasis-M	2.7
110992 Emphysema-F	12.3	112420 Psoriasis-M	14.1
110993 Emphysema-F	7.2	112425 Match Control Psoriasis-M	6.7
110994 Emphysema-F	4.6	104689 (MF) OA Bone-Backus	21.6
110995 Emphysema-F	20.3	104690 (MF) Adj "Normal" Bone-Backus	21.8
110996 Emphysema-F	7.1	104691 (MF) OA Synovium-Backus	14.1
110997 Asthma-M	2.5	104692 (BA) OA Cartilage-Backus	53.6
111001 Asthma-F	6.7	104694 (BA) OA Bone-Backus	14.8
111002 Asthma-F	5.7	104695 (BA) Adj "Normal" Bone-Backus	28.7
111003 Atopic Asthma-F	11.0	104696 (BA) OA Synovium-Backus	15.8
111004 Atopic Asthma-F	13.3	104700 (SS) OA Bone-Backus	11.6
111005 Atopic Asthma-F	12.2	104701 (SS) Adj "Normal" Bone-Backus	12.7
111006 Atopic Asthma-F	2.6	104702 (SS) OA Synovium-Backus	27.5
111417 Allergy-M	7.6	117093 OA Cartilage Rep7	6.3
112347 Allergy-M	0.0	112672 OA Bone5	6.0
112349 Normal Lung-F	0.0	112673 OA Synovium5	1.4
112357 Normal Lung-F	19.9	112674 OA Synovial Fluid cells5	3.0
112354 Normal Lung-M	4.0	117100 OA Cartilage Rep14	4.0
112374 Crohns-F	2.7	112756 OA Bone9	100.0
112389 Match Control Crohns-F	9.3	112757 OA Synovium9	0.9
112375 Crohns-F	2.0	112758 OA Synovial Fluid Cells9	3.8
112732 Match Control Crohns-F	12.6	117125 RA Cartilage Rep2	9.0
112725 Crohns-M	0.3	113492 Bone2 RA	8.1
112387 Match Control Crohns-M	5.0	113493 Synovium2 RA	2.5
112378 Crohns-M	0.0	113494 Syn Fluid Cells RA	5.3
112390 Match Control Crohns-M	6.0	113499 Cartilage4 RA	6.7
112726 Crohns-M	9.9	113500 Bone4 RA	7.0
112731 Match Control Crohns-M	8.1	113501 Synovium4 RA	4.4
112380 Ulcer Col-F	6.0	113502 Syn Fluid Cells4 RA	3.2
112734 Match Control Ulcer Col-F	21.0	113495 Cartilage3 RA	6.3
112384 Ulcer Col-F	14.1	113496 Bone3 RA	8.4
112737 Match Control Ulcer Col-F	3.4	113497 Synovium3 RA	5.1
112386 Ulcer Col-F	3.4	113498 Syn Fluid Cells3 RA	7.9
112738 Match Control Ulcer Col-F	18.0	117106 Normal Cartilage Rep20	8.7
112381 Ulcer Col-M	0.0	113663 Bone3 Normal	0.0
112735 Match Control Ulcer Col-M	0.5	113664 Synovium3 Normal	0.0

112382 Ulcer Col-M	7.1	113665 Syn Fluid Cells3 Normal	0.6
112394 Match Control Ulcer Col-M	1.6	117107 Normal Cartilage Rep22	1.8
112383 Ulcer Col-M	13.1	113667 Bone4 Normal	2.4
112736 Match Control Ulcer Col-M	3.8	113668 Synovium4 Normal	1.7
112423 Psoriasis-F	6.3	113669 Syn Fluid Cells4 Normal	3.9

Table AUD. CNS_neurodegeneration_v1.0

5

Tissue Name	Rel. Exp.(%) Ag4075, Run 214294982	Tissue Name	Rel. Exp.(%) Ag4075, Run 214294982
AD 1 Hippo	11.0	Control (Path) 3 Temporal Ctx	1.0
AD 2 Hippo	8.4	Control (Path) 4 Temporal Ctx	1.7
AD 3 Hippo	8.0	AD 1 Occipital Ctx	6.5
AD 4 Hippo	2.9	AD 2 Occipital Ctx (Missing)	0.0
AD 5 Hippo	16.8	AD 3 Occipital Ctx	1.3
AD 6 Hippo	100.0	AD 4 Occipital Ctx	3.6
Control 2 Hippo	19.6	AD 5 Occipital Ctx	11.9
Control 4 Hippo	17.6	AD 6 Occipital Ctx	6.5
Control (Path) 3 Hippo	3.0	Control 1 Occipital Ctx	5.6
AD 1 Temporal Ctx	6.3	Control 2 Occipital Ctx	10.4
AD 2 Temporal Ctx	14.1	Control 3 Occipital Ctx	6.0
AD 3 Temporal Ctx	4.2	Control 4 Occipital Ctx	2.9
AD 4 Temporal Ctx	7.5	Control (Path) 1 Occipital Ctx	3.3
AD 5 Inf Temporal Ctx	8.9	Control (Path) 2 Occipital Ctx	0.5
AD 5 Sup Temporal Ctx	24.5	Control (Path) 3 Occipital Ctx	1.6
AD 6 Inf Temporal Ctx	78.5	Control (Path) 4 Occipital Ctx	0.4
AD 6 Sup Temporal Ctx	56.6	Control 1 Parietal Ctx	5.9
Control 1 Temporal Ctx	2.3	Control 2 Parietal Ctx	9.9
Control 2 Temporal Ctx	12.1	Control 3 Parietal Ctx	6.0
Control 3 Temporal Ctx	7.7	Control (Path) 1 Parietal Ctx	3.6
Control 3 Temporal Ctx	3.1	Control (Path) 2 Parietal Ctx	1.1
Control (Path) 1 Temporal Ctx	4.6	Control (Path) 3 Parietal Ctx	2.2
Control (Path) 2 Temporal Ctx	1.8	Control (Path) 4 Parietal Ctx	3.4

Table AUE. General screening panel v1.4

Tissue Name	Rel. Exp.(%) Ag4075, Run 212696066	Rel. Exp.(%) Ag4075, Run 218525356	Tissue Name	Rel. Exp.(%) Ag4075, Run 212696066	Rel. Exp.(%) Ag4075, Run 218525356
Adipose	0.0	1.3	Renal ca. TK-10	9.7	14.8
Melanoma* Hs688(A).T	14.4	23.2	Bladder	1.0	1.8
Melanoma* Hs688(B).T	19.1	29.9	Gastric ca. (liver met.) NCI-N87	41.5	42.0
Melanoma* M14	9.5	12.7	Gastric ca. KATO III	25.5	22.8
Melanoma* LOXIMVI	8.1	12.9	Colon ca. SW-948	4.4	5.6
Melanoma* SK-MEL-5	5.9	14.2	Colon ca. SW480	100.0	100.0
Squamous cell carcinoma SCC-4	5.1	10.2	Colon ca.* (SW480 met) SW620	41.5	50.0
Testis Pool	1.4	1.9	Colon ca. HT29	10.2	13.6
Prostate ca.* (bone met) PC-3	9.5	13.6	Colon ca. HCT-116	13.0	20.9
Prostate Pool	1.1	1.5	Colon ca. CaCo-2	12.0	14.5
Placenta	1.1	1.3	Colon cancer tissue	5.0	8.4
Uterus Pool	0.1	0.2	Colon ca. SW1116	14.7	15.9
Ovarian ca. OVCAR-3	6.5	8.0	Colon ca. Colo-205	24.7	29.5
Ovarian ca. SK-OV-3	8.1	9.9	Colon ca. SW-48	3.6	4.7
Ovarian ca. OVCAR-4	9.2	16.4	Colon Pool	0.7	1.1
Ovarian ca. OVCAR-5	28.1	32.1	Small Intestine Pool	0.5	0.6
Ovarian ca. IGROV-1	23.0	33.2	Stomach Pool	0.8	0.8
Ovarian ca. OVCAR-8	10.3	16.4	Bone Marrow Pool	0.2	0.4
Ovary	0.5	0.8	Fetal Heart	0.1	0.1
Breast ca. MCF-7	15.7	17.2	Heart Pool	0.2	0.3
Breast ca. MDA-MB-231	10.4	15.6	Lymph Node Pool	1.2	1.0
Breast ca. BT 549	9.9	18.7	Fetal Skeletal Muscle	0.2	0.2
Breast ca. T47D	53.2	51.8	Skeletal Muscle Pool	0.2	0.3
Breast ca. MDA-N	4.7	6.3	Spleen Pool	0.7	0.5
Breast Pool	0.6	0.6	Thymus Pool	0.8	0.9

Trachea	3.6	5.3	CNS cancer (glio/astro) U87-MG	20.0	20.3
Lung	0.1	0.1	CNS cancer (glio/astro) U-118-MG	11.2	12.9
Fetal Lung	2.4	4.0	CNS cancer (neuro;met) SK-N-AS	6.9	8.9
Lung ca. NCI-N417	1.6	0.0	CNS cancer (astro) SF-539	9.3	12.0
Lung ca. LX-1	81.8	82.4	CNS cancer (astro) SNB-75	36.1	55.5
Lung ca. NCI-H146	0.4	0.8	CNS cancer (glio) SNB-19	30.1	37.6
Lung ca. SHP-77	6.8	8.5	CNS cancer (glio) SF-295	58.6	60.7
Lung ca. A549	9.8	15.8	Brain (Amygdala) Pool	0.0	0.1
Lung ca. NCI-H526	2.1	2.5	Brain (cerebellum)	0.1	0.2
Lung ca. NCI-H23	4.3	4.2	Brain (fetal)	0.2	0.3
Lung ca. NCI-H460	9.2	16.2	Brain (Hippocampus) Pool	0.1	0.1
Lung ca. HOP-62	4.4	4.5	Cerebral Cortex Pool	0.0	0.1
Lung ca. NCI-H522	9.5	10.0	Brain (Substantia nigra) Pool	0.1	0.1
Liver	0.0	0.1	Brain (Thalamus) Pool	0.0	0.1
Fetal Liver	2.9	4.3	Brain (whole)	0.2	0.2
Liver ca. HepG2	6.7	7.9	Spinal Cord Pool	0.2	0.3
Kidney Pool	1.1	1.2	Adrenal Gland	0.3	0.6
Fetal Kidney	0.3	0.5	Pituitary gland Pool	0.1	0.3
Renal ca. 786-0	5.1	9.5	Salivary Gland	3.0	2.8
Renal ca. A498	3.1	5.0	Thyroid (female)	0.1	0.1
Renal ca. ACHN	5.1	5.9	Pancreatic ca. CAPAN2	7.9	12.2
Renal ca. UO-31	2.6	4.2	Pancreas Pool	1.3	1.2

Table AUF. General screening panel v1.5

Tissue Name	Rel. Exp.(%) Ag4075, Run 228714883	Tissue Name	Rel. Exp.(%) Ag4075, Run 228714883
Adipose	1.0	Renal ca. TK-10	9.8
Melanoma* Hs688(A).T	18.0	Bladder	1.4
Melanoma* Hs688(B).T	17.4	Gastric ca. (liver met.) NCI-N87	35.4
Melanoma* M14	9.5	Gastric ca. KATO III	19.9
Melanoma* LOXIMVI	9.0	Colon ca. SW-948	4.4
Melanoma* SK-MEL-5	8.7	Colon ca. SW480	100.0
Squamous cell carcinoma SCC-4	5.8	Colon ca.* (SW480 met) SW620	32.8
Testis Pool	1.2	Colon ca. HT29	9.9
Prostate ca.* (bone met) PC-3	10.8	Colon ca. HCT-116	15.2
Prostate Pool	1.5	Colon ca. CaCo-2	11.1
Placenta	1.1	Colon cancer tissue	5.1
Uterus Pool	0.3	Colon ca. SW1116	7.2
Ovarian ca. OVCAR-3	6.2	Colon ca. Colo-205	23.7
Ovarian ca. SK-OV-3	7.5	Colon ca. SW-48	3.2
Ovarian ca. OVCAR-4	12.5	Colon Pool	0.7
Ovarian ca. OVCAR-5	20.2	Small Intestine Pool	0.4
Ovarian ca. IGROV-1	23.8	Stomach Pool	0.7
Ovarian ca. OVCAR-8	11.2	Bone Marrow Pool	0.2
Ovary	0.6	Fetal Heart	0.1
Breast ca. MCF-7	14.4	Heart Pool	0.2
Breast ca. MDA-MB-231	14.1	Lymph Node Pool	0.7
Breast ca. BT 549	8.4	Fetal Skeletal Muscle	0.2
Breast ca. T47D	2.1	Skeletal Muscle Pool	0.4
Breast ca. MDA-N	3.6	Spleen Pool	0.3
Breast Pool	0.5	Thymus Pool	0.5
Trachea	4.6	CNS cancer (glio/astro) U87-MG	12.5
Lung	0.1	CNS cancer (glio/astro) U-118-MG	8.5
Fetal Lung	2.6	CNS cancer (neuro;met) SK-N-AS	5.5
Lung ca. NCI-N417	1.9	CNS cancer (astro) SF-539	8.4
Lung ca. LX-1	81.8	CNS cancer (astro) SNB-75	13.1
Lung ca. NCI-H146	0.6	CNS cancer (glio) SNB-19	27.2
Lung ca. SHP-77	7.7	CNS cancer (glio) SF-295	53.2
Lung ca. A549	11.8	Brain (Amygdala) Pool	0.0
Lung ca. NCI-H526	2.1	Brain (cerebellum)	0.1
Lung ca. NCI-H23	3.5	Brain (fetal)	0.2
Lung ca. NCI-H460	8.8	Brain (Hippocampus) Pool	0.0
Lung ca. HOP-62	3.5	Cerebral Cortex Pool	0.1

Lung ca. NCI-H522	7.5	Brain (Substantia nigra) Pool	0.1
Liver	0.0	Brain (Thalamus) Pool	0.1
Fetal Liver	2.9	Brain (whole)	0.2
Liver ca. HepG2	6.2	Spinal Cord Pool	0.1
Kidney Pool	0.8	Adrenal Gland	0.4
Fetal Kidney	0.3	Pituitary gland Pool	0.2
Renal ca. 786-0	5.6	Salivary Gland	2.7
Renal ca. A498	3.4	Thyroid (female)	0.1
Renal ca. ACHN	4.9	Pancreatic ca. CAPAN2	9.7
Renal ca. UO-31	2.4	Pancreas Pool	0.8

Table AUG. Panel 3D

5

Tissue Name	Rel. Exp.() Ag4075, Run 186579982	Tissue Name	Rel. Exp.(%) Ag4075, Run 186579982
Daoy- Medulloblastoma	1.7	Ca Ski- Cervical epidermoid carcinoma (metastasis)	9.3
TE671- Medulloblastoma	1.3	ES-2- Ovarian clear cell carcinoma	4.2
D283 Med- Medulloblastoma	13.6	Ramos- Stimulated with PMA/ionomycin 6h	12.2
PFSK-1- Primitive Neuroectodermal	8.0	Ramos- Stimulated with PMA/ionomycin 14h	12.2
XF-498- CNS	5.1	MEG-01- Chronic myelogenous leukemia (megokaryoblast)	25.0
SNB-78- Glioma	12.9	Raji- Burkitt's lymphoma	2.4
SF-268- Glioblastoma	5.4	Daudi- Burkitt's lymphoma	5.0
T98G- Glioblastoma	7.9	U266- B-cell plasmacytoma	9.3
SK-N-SH- Neuroblastoma (metastasis)	4.4	CA46- Burkitt's lymphoma	2.6
SF-295- Glioblastoma	8.2	RL- non-Hodgkin's B-cell lymphoma	6.5
Cerebellum	0.1	JM1- pre-B-cell lymphoma	6.0
Cerebellum	0.1	Jurkat- T cell leukemia	7.6
NCI-H292- Mucoepidermoid lung carcinoma	12.0	TF-1- Erythroleukemia	17.6
DMS-114- Small cell lung cancer	3.0	HUT 78- T-cell lymphoma	4.9
DMS-79- Small cell lung cancer	92.0	U937- Histiocytic lymphoma	17.9
NCI-H146- Small cell lung cancer	1.6	KU-812- Myelogenous leukemia	15.4
NCI-H526- Small cell lung cancer	10.7	769-P- Clear cell renal carcinoma	5.8